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FILE COVERS 1907 - 9 Mar 2005 VOL 142 ISS 11 FILE LAST UPDATED: 8 Mar 2005 (20050308/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d his

(FILE 'HOME' ENTERED AT 12:52:20 ON 09 MAR 2005) SET COST OFF

FILE 'REGISTRY' ENTERED AT 12:52:25 ON 09 MAR 2005 E LACTOFERRIN

L1 224 S E3, E4, E8

L2 23 S E1, E2, E5-E7 NOT L1

FILE 'HCAPLUS' ENTERED AT 12:53:57 ON 09 MAR 2005

E LACTOFERRIN/CT

L3 3804 S E6-E10 E E6+ALL

L4 3828 S E4,E3

L5 4954 S LACTOFERRIN OR LACTOTRANSFERRIN

L6 349 S L1 OR L2

L7 5056 S L3-L6

L8 1 S LACTO() (FERRIN OR TRANSFERRIN OR TRANS FERRIN)

L9 5056 S L7, L8

E ATHEROSCLEROSIS/CT

L10 26376 S E3, E4

E E3+ALL

L11 5033 S E10-E13

L12 45035 S E9,E11,E12,E13/BI

E E8+ALL

L13 8095 S E8

L14 10796 S E8/BI

E E15+ALL

L15 8237 S E4

L16 32 S L9 AND L10-L15

E CARDIOVASCULAR/CT

E E5+ALL

L17 63843 S E3+NT

E E19+ALL

L18 245711 S E4,E3+NT

E E250+ALL

L19 375833 S E3+NT

```
E HEART DISEASE/CT
                E E4+ALL
                E E2+ALL
L20
          82991 S E8, E9, E7+NT
                E E92+ALL
L21
         216429 S E5, E4+NT
L22
           6523 S E10+OLD, NT
L23
            189 S L9 AND L17-L22
L24
             12 S L9 AND CARDIOVASCULAR(L) (DISEASE OR DISORDER OR DYSFUNCTION?)
L25
            194 S L16, L23, L24
     FILE 'REGISTRY' ENTERED AT 12:59:21 ON 09 MAR 2005
              1 S CHOLESTEROL/CN
L26
                E C-REACTIVE PROTEIN/CN
L27
              1 S E3
L28
            106 S C REACTIVE PROTEIN
     FILE 'HCAPLUS' ENTERED AT 13:00:01 ON 09 MAR 2005
L29
         107734 S L26 OR L27 OR L28
L30
             60 S L29 AND L9
            137 S (?CHOLESTER? OR CRP OR C REACTIVE(L) PROTEIN) AND L9
L31
             30 S TRIGLYCER? AND L9
L32
L33
             33 S ?VASCUL?(L)?INFLAM? AND L9
L34
              1 S ?VASCUL?(L)?SPASM? AND L9
              1 S BLOOD VESSEL+OLD, NT/CT (L) SPASM? AND L9
L35
              1 S BLOOD VESSEL, DISEASE+OLD, NT/CT (L) SPASM? AND L9
L36
             34 S PROTEIN?/CW,CT (L) C REACTIVE AND L9
L37
L38
              1 S BLOOD VESSEL, DISEASE+OLD, NT/CT (L) HYPERREACT? AND L9
L39
              1 S BLOOD VESSEL+OLD, NT/CT (L) HYPERREACT? AND L9
              6 S BLOOD VESSEL+OLD, NT/CT (L) SMOOTH MUSCL? AND L9
L40
              4 S BLOOD VESSEL, DISEASE+OLD, NT/CT (L) SMOOTH MUSCL? AND L9
1.41
              6 S INFLAMM?/CW,CT (L) VASCUL? AND L9
L42
             0 S INFLAMM?/CW,CT (L) PRO(L)CYTOKIN? AND L9
L43
             10 S INFLAMM?/CW,CT (L) CYTOKIN? AND L9
L44
             37 S CYTOKINE?/CW,CT (L) ?INFLAM? AND L9
L45
                E CYTOKINE/CT
             72 S E77+OLD,NT (L) ?INFLAM? AND L9
L46
             14 S BLOOD VESSEL, DISEASE+OLD, NT/CT (L) ENDOTHEL? AND L9
L47
L48
             39 S BLOOD VESSEL+OLD, NT/CT (L) ENDOTHEL? AND L9
L49
             12 S ENDOTHELIUM+OLD, NT/CT (L) VASCUL? AND L9
                E HYPERCHOLESTEROL/CT
L50
              7 S E5, E6 AND L9
                E E5+ALL
              0 S E5 AND L9
L51
                E HYPERTRIGLYCER/CT
                E E4+ALL
              4 S E4, E5 AND L9
L52
                E LOW DENSITY LIPOPROTEIN/CT
                E L DENSITY LIPOPROTEIN/CT
                E LIPOPROTEIN/CT
L53
             12 S E100-E109, E113, E114 AND L9
L54
             54 S E135-E146 AND L9
                E E51+ALL
             56 S E2+NT (L) (LOW OR VERY LOW) () (DENSITY OR D OR DEN) AND L9
1.55
L56
             12 S E2+NT (L) HIGH() (DENSITY OR D OR DEN) AND L9
L57
             19 S E2+NT (L) (LDL OR VLDL OR HDL OR VHDL) AND L9
L58
            401 S L30-L57, L25
              1 S US20040152623/PN OR WO2003-US38540/AP.PRN
L59
                E VARADHACHARY A/AU
L60
             19 S E3, E7
                E GLYNN P/AU
L61
             53 S E3-E9, E17-E19
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E WANG Y/AU

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L62
           2479 S E3, E40-E43
                E WANG YEN/AU
             11 S E3, E34
L63
L64
             13 S E50
                E ENGELMAYER J/AU
              9 S E4
L65
                E AGENNIX/AP,CS
                E AGENNIX/PA,CS
                E AGENIX/PA,CS
L66
             17 S E3-E21
L67
             10 S L59-L66 AND L58
L68
             10 S L67 AND L3-L25, L29-L67
L69
            341 S L58 AND (PD<=20021204 OR PRD<=20021204 OR AD<=20021204)
            106 S L69 AND (PHARMACEUT? OR PHARMACOL?)/SC,SX
L70
                E DRUG DELIVERY/CT
L71
              2 S E27-E31,E39 AND L70
L72
              0 S E53, E55, E58, E64, E70, E71 AND L70
L73
              0 S E89,E107 AND L70
L74
             50 S E6-E217 AND L70
                E E6+ALL
L75
              6 S E3-E5 AND L70
L76
             53 S E2+NT AND L70
L77
             53 S L71, L74-L76
L78
            106 S L70, L77
     FILE 'REGISTRY' ENTERED AT 13:36:48 ON 09 MAR 2005
     FILE 'REGISTRY' ENTERED AT 13:37:26 ON 09 MAR 2005
L79
             14 S 59-67-6 OR 943-45-3 OR 11041-12-6 OR 50925-79-6 OR 182815-43-
L80
            717 S (59-67-6 OR 943-45-3 OR 11041-12-6 OR 50925-79-6 OR 182815-43
L81
              1 S 9028-35-7
     FILE 'HCAPLUS' ENTERED AT 13:38:11 ON 09 MAR 2005
              6 S L79, L80, L81 AND L78
L82
L83
              8 S L79, L80, L81 AND L69
L84
              1 S BILE ACID (L) SEQUESTR? AND L69, L78
              7 S L82-L84, L78 AND ?ATHEROSCLERO?
L85
L86
             23 S L69 AND ?ATHEROSCLERO?
L87
             23 S L85, L86
                SEL DN AN 1 7 12
              3 S L87 AND E1-E7
L88
L89
             99 S L78 NOT L87
                SEL DN AN 8 19 21 68 89
                DEL SEL
                SEL DN AN 8 19 21 68 96
              5 S L89 AND E1-E15
L90
L91
             16 S L88, L90, L68
L92
             16 S L91 AND L3-L25, L29-L78, L82-L91
     FILE 'HCAPLUS' ENTERED AT 13:53:44 ON 09 MAR 2005
=> d all hitstr tot 192
    ANSWER 1 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
1.92
AN
     2005:77958 HCAPLUS
DN
     142:175360
ED
     Entered STN: 28 Jan 2005
     Lactoferrin as an adjuvant in cancer vaccines
TI
     Varadhachary, Atul; Pericle, Federica
IN
PA
     Agennix Incorporated, USA
     U.S. Pat. Appl. Publ., 22 pp., which
SO
     CODEN: USXXCO
DT
     Patent
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LA
    English
IC
    ICM A61K039-00
    ICS A61K038-40
NCL 424185100; 514006000
    15-2 (Immunochemistry)
FAN.CNT 1
    PATENT NO.
                      KIND
                              DATE
                                        APPLICATION NO.
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                              -----
                                         -----
                                                                -----
    US 2005019342
                       A1
                              20050127
                                        US 2004-862213
                                                             20040607
PRAI US 2003-476318P
                       P
                              20030606
    US 2003-498236P
                       P
                              20030827
CLASS
PATENT NO.
               CLASS PATENT FAMILY CLASSIFICATION CODES
 US 2005019342 ICM
                      A61K039-00
                ICS
                      A61K038-40
               NCL
                      424185100; 514006000
    The present invention relates to methods of treating cancer by
AB
    administering a composition of lactoferrin (LF) in combination with
    cancer vaccines. The examples describe the oral LF inhibition of
    Her-2/Neu+ transplantable carcinoma (TUBO) in mouse model; evaluation of
    recombinant human (rh) LF as adjuvant of p185 DNA vaccine in the
    prevention and treatment of TUBO in mice; oral LF in inhibition of
    spontaneous carcinomas; oral LF and DNA vaccination in inhibition of
    spontaneous carcinomas; oral LF in combination with tumor cell vaccination
    in inhibition of spontaneous carcinomas; and oral administration of hLF in
    combination with a cancer vaccine in patients.
ST
    lactoferrin adjuvant cancer vaccine
IT
    Leukemia
       (acute lymphocytic; lactoferrin as adjuvant in cancer
       vaccines)
IT
    Leukemia
       (acute myelogenous; lactoferrin as adjuvant in cancer
       vaccines)
IT
    Immunostimulants
       (adjuvants; lactoferrin as adjuvant in cancer vaccines)
IT
    Neuroglia, neoplasm
       (astrocytoma; lactoferrin as adjuvant in cancer vaccines)
IT
    Therapy
       (biotherapy; lactoferrin as adjuvant in cancer vaccines in
       combination with)
ΙT
    Bos taurus
       (bovine or human lactoferrin as adjuvant in cancer vaccines)
IT
    Drug delivery systems
       (carriers; lactoferrin as adjuvant in cancer vaccines)
    Uterus, neoplasm
IT
       (cervix; lactoferrin as adjuvant in cancer vaccines)
IT
    Leukemia
       (chronic lymphocytic; lactoferrin as adjuvant in cancer
       vaccines)
IT
    Leukemia
       (chronic myelocytic; lactoferrin as adjuvant in cancer
       vaccines)
IT
    Leukemia
       (chronic myelomonocytic leukemia; lactoferrin as adjuvant in
       cancer vaccines)
IT
    Intestine, neoplasm
       (colon; lactoferrin as adjuvant in cancer vaccines)
ΙT
    Neuroglia, neoplasm
       (glioblastoma; lactoferrin as adjuvant in cancer vaccines)
IT
    Mouth, neoplasm
       (gum; lactoferrin as adjuvant in cancer vaccines)
IT
    Neoplasm
```

```
(hematol.; lactoferrin as adjuvant in cancer vaccines)
IT
     Cytokines
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (immunomodulatory; lactoferrin as adjuvant in cancer vaccines
        containing)
IT
     Antitumor agents
     Bladder, neoplasm
     Bone, neoplasm
     Brain, neoplasm
     Combination chemotherapy
     Digestive tract, neoplasm
     Genetic vectors
     Head, neoplasm
     Human
     Immunotherapy
     Kidney, neoplasm
     Leukemia
     Lymphoma
     Mammary gland, neoplasm
     Melanoma
     Multiple myeloma
     Myelodysplastic syndromes
     Ovary, neoplasm
     Pancreas, neoplasm
     Prostate gland, neoplasm
     Sarcoma
     Testis, neoplasm
     Tongue, neoplasm
        (lactoferrin as adjuvant in cancer vaccines)
IT
     Lactoferrins
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lactoferrin as adjuvant in cancer vaccines)
ΙT
     Nucleic acids
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lactoferrin as adjuvant in cancer vaccines containing)
ΙT
     Chemotherapy
     Radiotherapy
     Surgery
        (lactoferrin as adjuvant in cancer vaccines in combination
        with)
IT
     Antigen-presenting cell
     CD4-positive T cell
     CD8-positive T cell
     Dendritic cell
     Hematopoietic precursor cell
     T cell (lymphocyte)
        (lactoferrin as adjuvant in cancer vaccines in relation to
        activation of)
IT
     Interleukin 18
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lactoferrin as adjuvant in cancer vaccines in relation to
        formation of)
IT
     neu (receptor)
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (lactoferrin as vaccine adjuvant in Neu-expressing cancers)
IT
     Chemokines
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (macrophage inflammatory protein 3\alpha;
```

lactoferrin as adjuvant in cancer vaccines in relation to formation of) IT Mesothelium, neoplasm (mesothelioma; lactoferrin as adjuvant in cancer vaccines) IT Lymphocyte (natural killer cell; lactoferrin as adjuvant in cancer vaccines in relation to activation of) IT Neoplasm (neck; lactoferrin as adjuvant in cancer vaccines) IT Astrocyte (neoplasm, astrocytoma; lactoferrin as adjuvant in cancer vaccines) IT Neck, anatomical (neoplasm; lactoferrin as adjuvant in cancer vaccines) IT Nerve, neoplasm (neuroblastoma; lactoferrin as adjuvant in cancer vaccines) IT Lung, neoplasm (non-small-cell carcinoma; lactoferrin as adjuvant in cancer vaccines) Drug delivery systems IT (oral; lactoferrin as adjuvant in cancer vaccines) IT Antiqens RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (p185; lactoferrin as adjuvant in cancer vaccines containing DNA encoding) IT Drug delivery systems (parenterals; lactoferrin as adjuvant in cancer vaccines) IT Carcinoma (pulmonary non-small-cell; lactoferrin as adjuvant in cancer vaccines) Carcinoma IT (pulmonary small-cell; lactoferrin as adjuvant in cancer vaccines) IT Eye, neoplasm (retinoblastoma; lactoferrin as adjuvant in cancer vaccines) IT Lung, neoplasm (small-cell carcinoma; lactoferrin as adjuvant in cancer vaccines) IT Carcinoma (squamous cell; lactoferrin as adjuvant in cancer vaccines) IT Drug delivery systems (topical; lactoferrin as adjuvant in cancer vaccines) IT Antigens RL: BSU (Biological study, unclassified); BIOL (Biological study) (tumor-associated; lactoferrin as adjuvant in DNA cancer vaccines promoting recognition of) IT Vaccines (tumor; lactoferrin as adjuvant in cancer vaccines) IT DNA RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (vaccine; lactoferrin as adjuvant in cancer vaccines containing) IT Antitumor agents (vaccines; lactoferrin as adjuvant in cancer vaccines) 56-40-6, Glycine, biological studies IT RL: BSU (Biological study, unclassified); BIOL (Biological study) (N-terminal glycine absence in lactoferrin variant) 83869-56-1, Granulocyte-macrophage colony-stimulating factor IT RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (lactoferrin as adjuvant in cancer vaccines in relation to formation of)

```
IT
    832811-61-7
                  832811-62-8
                                832811-63-9 832811-64-0
    RL: PRP (Properties)
        (unclaimed nucleotide sequence; lactoferrin as an adjuvant in
        cancer vaccines)
IT
    151812-50-9
                  154427-28-8
                               160212-35-1
                                             163816-02-2
                                                           204380-35-8
    473461-56-2
    RL: PRP (Properties)
        (unclaimed sequence; lactoferrin as an adjuvant in cancer
       vaccines)
L92
    ANSWER 2 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    2004:1033558 HCAPLUS
DN
    141:420455
ED
    Entered STN: 02 Dec 2004
    Compositions comprising recombinant lactoferrin and its variants
TI
    in the treatment of diabetes mellitus
IN
    Engelmayer, Jose; Varadhachary, Atul
PA
    Agennix Incorporated, USA
    PCT Int. Appl., 32 pp.
so
    CODEN: PIXXD2
DT
    Patent
    English
LA
IC
    ICM A61K
CC
    1-10 (Pharmacology)
FAN.CNT 1
    PATENT NO.
                                        APPLICATION NO.
                       KIND DATE
                                                               DATE
     ______
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                              -----
                                          _______
PΤ
    WO 2004103285
                        A2
                              20041202 WO 2004-US14985
                                                               20040513
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
            NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
            TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
            AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
            EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
            SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
            SN, TD, TG
    US 2005004006
                        A1
                              20050106
                                          US 2004-844865
                                                                20040513
PRAI US 2003-470549P
                        Ρ
                              20030514
CLASS
PATENT NO.
               CLASS PATENT FAMILY CLASSIFICATION CODES
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WO 2004103285 ICM
                       A61K
AΒ
    The present invention relates to methods of using a composition of
    lactoferrin for the treatment of diabetes mellitus as manifested
    by a reduction in the levels of serum glucose, blood pressure, obesity, or
    glycosylated Hb (HbA1c).
ST
    human recombinant lactoferrin variant antidiabetic
    hyperglycemia; recombinant lactoferrin antiobesity
    antihypertensive
IT
    Drug delivery systems
        (carriers; compns. comprising recombinant lactoferrin and its
       variants in treatment of diabetes mellitus)
IT
    Antidiabetic agents
      Antihypertensives
    Antiobesity agents
    Chelating agents
    Diabetes mellitus
    Hyperglycemia
      Hypertension
    Obesity
```

(compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) Lactoferrins IT RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Hemoglobins RL: BSU (Biological study, unclassified); BIOL (Biological study) (glycohemoglobins; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Drug delivery systems (injections, i.m.; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Drug delivery systems (injections, i.v.; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Drug delivery systems (injections, s.c.; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) Diabetes mellitus IT (insulin-dependent; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) TT Body weight (lactoferrin composition reducing; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Bos taurus Human Mammalia (lactoferrin; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) TΤ Intestine (large, lactoferrin releasing in; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) Diabetes mellitus IT(non-insulin-dependent; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Drug delivery systems (oral; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Drug delivery systems (parenterals; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Intestine (small, lactoferrin releasing in; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) IT Drug delivery systems (transdermal; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) ΙT 60-00-4, EDTA, biological studies 67-42-5, EGTA 150-39-0, HEDTA 85233-19-8, BAPTA RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (as metal chelator; compns. comprising recombinant lactoferrin and its variants in treatment of diabetes mellitus) 50-99-7, D-Glucose, biological studies IT RL: BSU (Biological study, unclassified); BIOL (Biological study)

variants in treatment of diabetes mellitus)

(blood; compns. comprising recombinant lactoferrin and its

and its variants in treatment of diabetes mellitus)

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L92
    ANSWER 3 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    2004:531382 HCAPLUS
DN
    141:47367
    Entered STN: 02 Jul 2004
ED
    Lactoferrin for the reduction of pain
TI
IN
    Varadhachary, Atul; Petrak, Karel
    Agennix Incorporated, USA
PA
SO
    PCT Int. Appl., 30 pp.
    CODEN: PIXXD2
DT
    Patent
LΑ
    English
IC
    ICM A61K038-40
    ICS C07K014-00
CC
    1-11 (Pharmacology)
    Section cross-reference(s): 63
FAN.CNT 1
                                        APPLICATION NO.
    PATENT NO.
                      KIND DATE
                                                              DATE
                              -----
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                                         ______
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    WO 2004054608
                       A2
                              20040701
                                         WO 2003-US39358
                                                               20031211
PΤ
    WO 2004054608
                       A3
                              20040805
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
            NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
            TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
            BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
            TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                       US 2003-733621
                              20040805
    US 2004151784
                        A1
                                                              20031211
PRAI US 2002-432937P
                        Р
                              20021212
    US 2003-498248P
                        Ρ
                              20030827
CLASS
               CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
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                ICM
WO 2004054608
                      A61K038-40
                ICS
                      C07K014-00
US 2004151784 ECLA A61K038/40; A61K038/40+M
    The invention discloses methods of using lactoferrin to reduce
AΒ
    pain in conditions associated with severe or intractable pain by
    administering a composition of lactoferrin either alone or in
    combination with other therapy for pain.
    lactoferrin pain treatment
ST
IT
    Tumor necrosis factors a
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (TNF-\alpha; lactoferrin for reduction of pain. and use with
       other therapeutic means)
IT
    Pain
        (acute; lactoferrin for reduction of pain. and use with other
       therapeutic means)
IT
    Acupuncture
        (and acupressure; lactoferrin for reduction of pain. and use with
       other therapeutic means)
IT
    Lactoferrins
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (and variants; lactoferrin for reduction of pain. and use with
       other therapeutic means)
    Bos taurus
IT
        (bovine lactoferrin; lactoferrin for reduction of pain.
```

and use with other therapeutic means) ITMetals, biological studies RL: BSU (Biological study, unclassified); BIOL (Biological study) (chelators for; lactoferrin for reduction of pain. and use with other therapeutic means) IT Therapy (chiropractic; lactoferrin for reduction of pain. and use with other therapeutic means) IT (chronic; lactoferrin for reduction of pain. and use with other therapeutic means) IT Drug delivery systems (delayed release; lactoferrin for reduction of pain. and use with other therapeutic means) IT Anesthesia (general; lactoferrin for reduction of pain. and use with other therapeutic means) IT Anesthetics (i.v.; lactoferrin for reduction of pain. and use with other therapeutic means) IT Analgesics Antacids Antidepressants Chelating agents Drug delivery systems Human Pain (lactoferrin for reduction of pain. and use with other therapeutic means) TΤ Cytokines RL: BSU (Biological study, unclassified); BIOL (Biological study) (lactoferrin for reduction of pain. and use with other therapeutic means) TT Opioids RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (lactoferrin for reduction of pain. and use with other therapeutic means) TT Intestine (large, lactoferrin release in; lactoferrin for reduction of pain. and use with other therapeutic means) IT Anesthesia (local; lactoferrin for reduction of pain. and use with other therapeutic means) IT Therapy (non-pharmacol. pain management techniques; lactoferrin for reduction of pain. and use with other therapeutic means) IT Anti-inflammatory agents (nonsteroidal; lactoferrin for reduction of pain. and use with other therapeutic means) IT Drug delivery systems (opioid pump; lactoferrin for reduction of pain. and use with other therapeutic means) ΤТ Drug delivery systems (oral; lactoferrin for reduction of pain. and use with other therapeutic means) IT Drug delivery systems (parenterals; lactoferrin for reduction of pain. and use with other therapeutic means) TT Cytokines RL: BSU (Biological study, unclassified); BIOL (Biological study) (proinflammatory; lactoferrin for reduction of pain.

and use with other therapeutic means)

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IT
    Anesthesia
        (regional; lactoferrin for reduction of pain. and use with other
       therapeutic means)
IT
    Intestine
        (small, lactoferrin release in; lactoferrin for
       reduction of pain. and use with other therapeutic means)
IT
        (spinal; lactoferrin for reduction of pain. and use with other
       therapeutic means)
    Drug delivery systems
IT
        (topical; lactoferrin for reduction of pain. and use with other
       therapeutic means)
IT
    56-40-6, Glycine, properties
    RL: PRP (Properties)
        (amino-terminal; lactoferrin for reduction of pain. and use with
       other therapeutic means)
    60-00-4, EDTA, biological studies 67-42-5, EGTA
IT
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lactoferrin for reduction of pain. and use with other
       therapeutic means)
    ANSWER 4 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
L92
AN
    2004:513498 HCAPLUS
DN
    141:47315
    Entered STN: 25 Jun 2004
ED
TI
    Lactoferrin as an agent in the prevention of organ transplant
    rejection and graft-versus-host-disease
IN
    Varadhachary, Atul; Pericle, Federica
PA
    Agennix Incorporated, USA
SO
    PCT Int. Appl., 38 pp.
    CODEN: PIXXD2
DT
    Patent
    English
LA
IC
    ICM A61K
CC
    1-7 (Pharmacology)
    Section cross-reference(s): 2, 63
FAN.CNT 1
    PATENT NO.
                                        APPLICATION NO.
                       KIND
                              DATE
                                                               DATE
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    WO 2004052305
                       A2
                                        WO 2003-US39265
PΤ
                              20040624
                                                               20031210
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            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
            NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
            TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
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            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
            TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                         US 2003-732429
    US 2004176276
                       A1
                              20040909
                                                               20031210
PRAI US 2002-432113P
                        P
                              20021210
    US 2003-498338P
                              20030827
                        Р
CLASS
              CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
 ______
WO 2004052305 ICM A61K
    The present invention relates to methods of using lactoferrin
AB
    (LF) to treat, prevent or reduce the incidence of organ transplant
    rejection and graft-vs.-host-disease. More particularly, the present
    invention relates to methods of reducing an immune response against
    miss-matched transplanted organs such as kidney, heart, lung, liver,
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0:

pancreas and stem cells by administering a composition of lactoferrin to the recipient patients. In addition, this invention relates to the treatment of bone marrow transplant (BMT) donors with lactoferrin to attenuate the development of graft-vs.-host-disease in the recipients. Moreover, this invention relates to the treatment of xenograft organ donors with lactoferrin to attenuate the development of graft rejection in the recipients. lactoferrin immunomodulator transplant rejection xenograft ST IT CD3 (antigen) CD4 (antigen) CD8 (antigen) RL: BSU (Biological study, unclassified); BIOL (Biological study) (T-lymphocytes expressing; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) Drug delivery systems TΤ (carriers; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) TΤ Drug delivery systems (delayed release; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) TΤ Transplant and Transplantation (graft-vs.-host reaction; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) Transplant and Transplantation IT (heart; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) IT Transplant and Transplantation (kidney; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) TΤ Bos taurus Human Immune tolerance Immunosuppressants Molecular cloning Stem cell Transplant and Transplantation Transplant rejection (lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) IT Lactoferrins RL: BPN (Biosynthetic preparation); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) IT Interleukin 18 RL: BSU (Biological study, unclassified); BIOL (Biological study) (lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) IT Intestine (large, lactoferrin release in; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) IT Transplant and Transplantation Transplant and Transplantation (liver; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) Transplant and Transplantation TT Transplant and Transplantation (lung; lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease) IT Chemokines RL: BSU (Biological study, unclassified); BIOL (Biological study)

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(macrophage inflammatory protein 3\alpha;
        lactoferrin as an agent in the prevention of organ transplant
        rejection and graft-vs.-host-disease)
IT
     Immunity
        (mucosal; lactoferrin as an agent in the prevention of organ
        transplant rejection and graft-vs.-host-disease)
IT
     Lymphocyte
        (natural killer cell, regulation of; lactoferrin as an agent
        in the prevention of organ transplant rejection and
        graft-vs.-host-disease)
     Drug delivery systems
IT
        (oral; lactoferrin as an agent in the prevention of organ
        transplant rejection and graft-vs.-host-disease)
     Transplant and Transplantation
IT
        (pancreas; lactoferrin as an agent in the prevention of organ
        transplant rejection and graft-vs.-host-disease)
IT
     Drug delivery systems
        (parenterals; lactoferrin as an agent in the prevention of
        organ transplant rejection and graft-vs.-host-disease)
IT
     Cytokines
     RL: ADV (Adverse effect, including toxicity); BIOL (Biological study)
        (proinflammatory; lactoferrin as an agent in the
        prevention of organ transplant rejection and graft-vs.-host-disease)
IT
     Antigen-presenting cell
     B cell (lymphocyte)
     Macrophage
     Polymorphonuclear leukocyte
     T cell (lymphocyte)
        (regulation of; lactoferrin as an agent in the prevention of
        organ transplant rejection and graft-vs.-host-disease)
ŤΤ
     Intestine
        (small, lactoferrin release in; lactoferrin as an
        agent in the prevention of organ transplant rejection and
        graft-vs.-host-disease)
IT
    Bone marrow
        (stem cells of; lactoferrin as an agent in the prevention of
        organ transplant rejection and graft-vs.-host-disease)
IT
     Immunity
        (systemic; lactoferrin as an agent in the prevention of organ
        transplant rejection and graft-vs.-host-disease)
IT
     Bone marrow
        (toxicity, stem cells of; lactoferrin as an agent in the
        prevention of organ transplant rejection and graft-vs.-host-disease)
TT
    Heart
    Kidney
    Liver
    Liver
    Lung
     Lung .
     Pancreas
        (transplant; lactoferrin as an agent in the prevention of
        organ transplant rejection and graft-vs.-host-disease)
     Transplant and Transplantation
IT
        (xenotransplant; lactoferrin as an agent in the prevention of
        organ transplant rejection and graft-vs.-host-disease)
     53-03-2, Prednisone 446-86-6, Azathioprine
IT
                                                   59865-13-3, Cyclosporine
     104987-11-3, Tacrolimus
                              128794-94-5, Mycophenolate mofetil
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lactoferrin as an agent in the prevention of organ
        transplant rejection and graft-vs.-host-disease)
IT
     60-00-4, Edta, biological studies
                                        67-42-5, Egta
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
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(lactoferrin as an agent in the prevention of organ transplant rejection and graft-vs.-host-disease)

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L92
    ANSWER 5 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
    2004:513478 HCAPLUS
AN
    141:17581
DN
ED
    Entered STN: 25 Jun 2004
    Oral lactoferrin for the treatment of sepsis
TI
IN
    Varadhachary, Atul; Petrak, Karel
PA
    Agennix Incorporated, USA
SO
    PCT Int. Appl., 44 pp.
    CODEN: PIXXD2
DТ
    Patent
LA
    English
IC
    ICM A61K
CC
    1-5 (Pharmacology)
    Section cross-reference(s): 63
FAN.CNT 1
    PATENT NO.
                      KIND DATE
                                        APPLICATION NO.
                                                              DATE
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    WO 2004052281
                                                               -----
                       A2
                              20040624
PΙ
                                        WO 2003-US38621
                                                              20031205
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    WO 2004052281
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            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
            NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
            TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
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    US 2004152624
                                       US 2003-728521
                       A1
                              20040805
                                                              20031205
PRAI US 2002-431393P
                        Ρ
                              20021206
    US 2003-498327P
                        P
                              20030827
CLASS
PATENT NO.
              CLASS PATENT FAMILY CLASSIFICATION CODES
 WO 2004052281 ICM A61K
    The invention relates to methods of treating prophylactically or
AB
    therapeutically bacteremia, sepsis, septic shock or related conditions
    such as ARDS by administering orally a composition of lactoferrin
    alone or in combination with standard therapies or metal chelators to prevent
    or treat the consequences of bacteria-induced systemic inflammatory
    response syndrome. In particular it is claimed that the therapeutic use
    of recombinant human lactoferrin alone or in combination with
    metal chelators or other therapeutic interventions decreases the mortality
    due to bacteremia, sepsis, septic shock or related conditions such as
    ARDS.
ST
    lactoferrin oral metal chelator bacteremia sepsis septic shock
    ARDS
IT
    Lung, disease
       (acute injury; oral lactoferrin for treatment of sepsis)
IT
    Injury
       (acute pulmonary; oral lactoferrin for treatment of sepsis)
IT
    Respiratory distress syndrome
       (acute; oral lactoferrin for treatment of sepsis)
IT
    Drug delivery systems
       (capsules, enteric-coated; oral lactoferrin for treatment of
       sepsis)
IT
    Drug delivery systems
       (carriers; oral lactoferrin for treatment of sepsis)
IT
    Organ, animal, disease
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(failure; oral lactoferrin for treatment of sepsis)
ΙT
     Drug delivery systems
        (infusions, nasogastric; oral lactoferrin for treatment of
        sepsis)
IT
     Drug delivery systems
        (injections, i.v.; oral lactoferrin for treatment of sepsis)
IT
     Immunity
        (mucosal; oral lactoferrin for treatment of sepsis)
IT
     Antacids
     Anti-inflammatory agents
     Antibiotics
     Bacteremia
     Blood
     Blood plasma
     Chelating agents
     Death
     Digestive tract
     Drug interactions
     Escherichia coli
     Eubacteria
     Haemophilus
     Human
     Immune system
     Immunomodulators
     Kidney
     Liver
     Lung
     Mucous membrane
     Pseudomonas
     Sepsis
     Spleen
     Staphylococcus
     Surfactants
        (oral lactoferrin for treatment of sepsis)
IT
     Interleukin 1
     Interleukin 10
     Interleukin 18
     Interleukin 2
     Interleukin 4
     Interleukin 5
     Interleukin 6
     Interleukin 8
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (oral lactoferrin for treatment of sepsis)
IT
     Lactoferrins
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (oral lactoferrin for treatment of sepsis)
IT
     Drug delivery systems
        (oral; oral lactoferrin for treatment of sepsis)
IT
     Cytokines
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (pro-inflammatory; oral lactoferrin for treatment
        of sepsis)
IT
     Shock (circulatory collapse)
        (septic; oral lactoferrin for treatment of sepsis)
IT
     Kidney
     Liver
     Lung
        (toxicity; oral lactoferrin for treatment of sepsis)
IT
     Medical goods
        (tubes, nasogastric; oral lactoferrin for treatment of
        sepsis)
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IT
    60-00-4, Ethylenediaminetetraacetic acid, biological studies 67-42-5,
    Ethylenebis(oxyethylenenitrilo)]tetraacetic acid 98530-76-8, Xigris
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (oral lactoferrin for treatment of sepsis)
L92
    ANSWER 6 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    2004:490705 HCAPLUS
    141:33800
DN
    Entered STN: 17 Jun 2004
ED
    Lactoferrin in the reduction of circulating cholesterol
ТT
     , vascular inflammation, atherosclerosis and
    cardiovascular disease
    Varadhachary, Atul; Glynn, Peter; Wang, Yenyun
IN
     ; Engelmayer, Jose
    Agennix Incorporated, USA
PA
    PCT Int. Appl., 38 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
    ICM A61K
TC
    1-8 (Pharmacology)
CC
FAN.CNT 1
    PATENT NO.
                      KIND DATE
                                        APPLICATION NO.
                                                              DATE
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    WO 2004050037
                       A2
                              20040617
                                       WO 2003-US38540
                                                              20031204 <--
PΙ
                       A3
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                              20040812
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            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC,
            LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
            NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
            TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
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            ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
            TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
    US 2004152623
                              20040805 US 2003-728275
                                                              20031204 <--
                    A1
                      P
PRAI US 2002-430867P
                              20021204
                                       <--
    US 2003-498337P
                       P
                              20030827
CLASS
               CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
 _____
WO 2004050037 ICM
                      A61K
    The invention discloses methods for using lactoferrin to reduce
    circulating levels of cholesterol and vascular
    inflammation in order to treat, prevent or reduce the incidence of
    atherosclerosis and cardiovascular disease.
ST
    lactoferrin hypocholesterolemic vascular
    inflammation atherosclerosis cardiovascular
    disease
TT
    Proteins
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (C-reactive; lactoferrin for reduction of
       cholesterol and vascular inflammation and
       treatment of atherosclerosis and cardiovascular
       disease)
TT
    Antiarteriosclerotics
        (antiatherosclerotics; lactoferrin for reduction of
       cholesterol and vascular inflammation and
       treatment of atherosclerosis and cardiovascular
       disease)
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IT

Sequestering agents

(bile acid; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease, and use with other agents) IT Bos taurus (bovine lactoferrin; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) Drug delivery systems IT (delayed release; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) ΙT Blood vessel (endothelium; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) IT Lipoproteins RL: BSU (Biological study, unclassified); BIOL (Biological study) (high-d.; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) Blood vessel IT (hyperreactivity; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) IT Drug delivery systems (injections, i.m.; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) IT Drug delivery systems (injections, i.p.; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) TΤ Drug delivery systems (injections, i.v.; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) TΤ Drug delivery systems (injections, s.c.; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) IT Drug delivery systems (intraarterial; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) TΤ Drug delivery systems (intramyocardial; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) Drug delivery systems TT (intrathecal; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of

atherosclerosis and cardiovascular disease)

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ΙT
    Anti-inflammatory agents
       Anticholesteremic agents
       Atherosclerosis
       Cardiovascular agents
       Cardiovascular system, disease
       Drug delivery systems
       Hypercholesterolemia
       Hypertriglyceridemia
     Hypolipemic agents
        (lactoferrin for reduction of cholesterol and
        vascular inflammation and treatment of
        atherosclerosis and cardiovascular disease)
IT
     Lactoferrins
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lactoferrin for reduction of cholesterol and
        vascular inflammation and treatment of
        atherosclerosis and cardiovascular disease)
IT
     Antacids
        (lactoferrin for reduction of cholesterol and
        vascular inflammation and treatment of
        atherosclerosis and cardiovascular disease,
        and use with other agents)
IT
     Intestine
        (large, lactoferrin release in; lactoferrin for
        reduction of cholesterol and vascular
        inflammation and treatment of atherosclerosis and
        cardiovascular disease)
TТ
     Lipoproteins
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (low-d.; lactoferrin for reduction of
        cholesterol and vascular inflammation and
        treatment of atherosclerosis and cardiovascular
        disease)
TТ
    Drug delivery systems
        (oral; lactoferrin for reduction of cholesterol
        and vascular inflammation and treatment of
        atherosclerosis and cardiovascular disease)
IT
     Drug delivery systems
        (parenterals; lactoferrin for reduction of
        cholesterol and vascular inflammation and
        treatment of atherosclerosis and cardiovascular
        disease)
TT
     Immunosuppressants
        (pro-inflammatory cytokine reduction; lactoferrin for
        reduction of cholesterol and vascular
        inflammation and treatment of atherosclerosis and
        cardiovascular disease)
TT
     Cytokines
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (proinflammatory; lactoferrin for reduction of
        cholesterol and vascular inflammation and
        treatment of atherosclerosis and cardiovascular
        disease)
TT
     Bile acids
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (sequestrants; lactoferrin for reduction of
        cholesterol and vascular inflammation and
        treatment of atherosclerosis and cardiovascular
        disease, and use with other agents)
     Intestine
IT
```

(small, lactoferrin release in; lactoferrin for

robinson - 10 / 728275 reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) IT Blood vessel (smooth muscle; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) ITBlood vessel, disease (spasm; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) IT Drug interactions (synergistic; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease, and use with other agents) Drug delivery systems IT (transendocardial; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) IT Drug delivery systems (transepicardial; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease) IT Biological transport (uptake, cholesterol absorption inhibitors; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease, and use with other agents) IT Cell proliferation Cytotoxic agents (vascular smooth muscle cell; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of

atherosclerosis and cardiovascular disease)

IT Endothelium

(vascular; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease)

IT Blood vessel, disease

Inflammation

(vasculitis; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease)

IT Lipoproteins

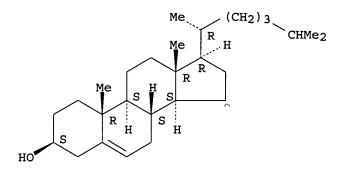
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(very-low-d.; lactoferrin for
reduction of cholesterol and vascular
inflammation and treatment of atherosclerosis and
cardiovascular disease)

IT 9028-35-7, HMG-CoA reductase

RL: BSU (Biological study, unclassified); BIOL (Biological study) (inhibitors; lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease, and use with other agents)

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IT
     57-88-5, Cholesterol, biological studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (lactoferrin for reduction of cholesterol and
        vascular inflammation and treatment of
        atherosclerosis and cardiovascular disease)
IT
     59-67-6, Nicotinic acid, biological studies 637-07-0,
     Clofibrate 943-45-3D, Fibric acid, derivs. 11041-12-6,
     Cholestyramine 25812-30-0, Gemfibrozil 49562-28-9,
     Fenofibrate 50925-79-6, Cholestipol 75330-75-5,
     Lovastatin 79902-63-9, Simvastatin 81093-37-0,
     Pravastatin 93957-54-1, Fluvastatin 134523-00-5,
     Atorvastatin 145599-86-6, Cerivastatin 182815-43-6,
     Colesevelam
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lactoferrin for reduction of cholesterol and
        vascular inflammation and treatment of
        atherosclerosis and cardiovascular disease,
        and use with other agents)
IT
     9028-35-7, HMG-CoA reductase
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (inhibitors; lactoferrin for reduction of cholesterol
        and vascular inflammation and treatment of
        atherosclerosis and cardiovascular disease,
        and use with other agents)
RN
     9028-35-7 HCAPLUS
CN
     Reductase, hydroxymethylglutaryl coenzyme A (reduced nicotinamide adenine
     dinucleotide phosphate) (9CI)
                                   (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     57-88-5, Cholesterol, biological studies
TT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (lactoferrin for reduction of cholesterol and
        vascular inflammation and treatment of
        atherosclerosis and cardiovascular disease)
RN
     57-88-5 HCAPLUS
     Cholest-5-en-3-ol (3β)- (9CI) (CA INDEX NAME)
CN
```

Absolute stereochemistry.



TT 59-67-6, Nicotinic acid, biological studies 637-07-0, Clofibrate 943-45-3D, Fibric acid, derivs. 11041-12-6, Cholestyramine 25812-30-0, Gemfibrozil 49562-28-9, Fenofibrate 50925-79-6, Cholestipol 75330-75-5, Lovastatin 79902-63-9, Simvastatin 81093-37-0, Pravastatin 93957-54-1, Fluvastatin 134523-00-5, Atorvastatin 145599-86-6, Cerivastatin 182815-43-6, Colesevelam RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(lactoferrin for reduction of cholesterol and vascular inflammation and treatment of atherosclerosis and cardiovascular disease, and use with other agents)

RN 59-67-6 HCAPLUS

CN 3-Pyridinecarboxylic acid (9CI) (CA INDEX NAME)

RN 637-07-0 HCAPLUS

CN Propanoic acid, 2-(4-chlorophenoxy)-2-methyl-, ethyl ester (9CI) (CA INDEX NAME)

RN 943-45-3 HCAPLUS

CN Propanoic acid, 2-methyl-2-phenoxy- (9CI) (CA INDEX NAME)

RN 11041-12-6 HCAPLUS

CN Cholestyramine (7CI, 8CI, 9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 25812-30-0 HCAPLUS

CN Pentanoic acid, 5-(2,5-dimethylphenoxy)-2,2-dimethyl- (9CI) (CA INDEX NAME)

RN 49562-28-9 HCAPLUS

CN Propanoic acid, 2-[4-(4-chlorobenzoyl)phenoxy]-2-methyl-, 1-methylethyl ester (9CI) (CA INDEX NAME)

RN 50925-79-6 HCAPLUS

CN Colestipol (9CI) (CA INDEX NAME)

*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***

RN 75330-75-5 HCAPLUS

CN Butanoic acid, 2-methyl-, (1S,3R,7S,8S,8aR)-1,2,3,7,8,8a-hexahydro-3,7-dimethyl-8-[2-[(2R,4R)-tetrahydro-4-hydroxy-6-oxo-2H-pyran-2-yl]ethyl]-1-naphthalenyl ester, (2S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 79902-63-9 HCAPLUS

CN Butanoic acid, 2,2-dimethyl-, (1S,3R,7S,8S,8aR)-1,2,3,7,8,8a-hexahydro-3,7-dimethyl-8-[2-[(2R,4R)-tetrahydro-4-hydroxy-6-oxo-2H-pyran-2-yl]ethyl]-1-naphthalenyl ester (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 81093-37-0 HCAPLUS

CN 1-Naphthaleneheptanoic acid, 1,2,6,7,8,8a-hexahydro- β , δ ,6-trihydroxy-2-methyl-8-[(2S)-2-methyl-1-oxobutoxy]-,

 $(\beta R, \delta R, 1S, 2S, 6S, 8S, 8aR) - (9CI)$ (CA INDEX NAME)

Absolute stereochemistry.

RN 93957-54-1 HCAPLUS

CN 6-Heptenoic acid, 7-[3-(4-fluorophenyl)-1-(1-methylethyl)-1H-indol-2-yl]-3,5-dihydroxy-, (3R,5S,6E)-rel-(9CI) (CA INDEX NAME)

Relative stereochemistry.
Double bond geometry as shown.

RN 134523-00-5 HCAPLUS

CN 1H-Pyrrole-1-heptanoic acid, 2-(4-fluorophenyl)- β , δ -dihydroxy-5-(1-methylethyl)-3-phenyl-4-[(phenylamino)carbonyl]-, (β R, δ R)-(9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 145599-86-6 HCAPLUS

CN 6-Heptenoic acid, 7-[4-(4-fluorophenyl)-5-(methoxymethyl)-2,6-bis(1-methylethyl)-3-pyridinyl]-3,5-dihydroxy-, (3R,5S,6E)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+). Double bond geometry as shown.

RN 182815-43-6 HCAPLUS

CN 1-Hexanaminium, N,N,N-trimethyl-6-(2-propenylamino)-, chloride, polymer with (chloromethyl)oxirane, 2-propen-1-amine and N-2-propenyl-1-decanamine (9CI) (CA INDEX NAME)

CM 1

CRN 182815-42-5 CMF C12 H27 N2 . Cl

 $H_2C = CH - CH_2 - NH - (CH_2)_6 - N + Me_3$

● Cl -

CM 2

CRN 92162-19-1 CMF C13 H27 N

 $H_2C = CH - CH_2 - NH - (CH_2)_9 - Me$

CM 3

CRN 107-11-9 CMF C3 H7 N

 $H_2C = CH - CH_2 - NH_2$

CM 4

CRN 106-89-8 CMF C3 H5 Cl O

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L92 ANSWER 7 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    2004:252366 HCAPLUS
DN
    140:276199
    Entered STN: 26 Mar 2004
ED
    Lactoferrin compositions and methods of wound treatment
TΙ
    Engelmayer, Jose; Varadhachary, Atul
IN
    Agennix Incorporated, USA
PA
    PCT Int. Appl., 62 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
    ICM A61K038-40
IC
    ICS C07K014-79
CC
    63-6 (Pharmaceuticals)
    Section cross-reference(s): 1
FAN.CNT 1
                                      APPLICATION NO.
    PATENT NO.
                     KIND DATE
    WO 2004024180 A1 20040325
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                             20040325 WO 2003-US29069
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                                                              20030916 <--
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            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE,
            GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK,
            LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ,
            OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM,
            TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
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            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                       A1 20040722 US 2003-663258
    US 2004142037
                                                              20030916 <--
PRAI US 2002-410981P
                        P
                              20020916 <--
CLASS
PATENT NO. CLASS PATENT FAMILY CLASSIFICATION CODES
 WO 2004024180 ICM
                      A61K038-40
                ICS
                      C07K014-79
US 2004142037
                ECLA
                      A61K038/40; A61K038/40+M; C07K014/79
    The present invention relates to lactoferrin compns. and methods
    of using the compns. to treat wounds. The compns. can be administered
    alone or in combination with other standard wound healing therapies.
    Lactoferrin enhances the local immune system and kills bacteria
    infecting the wound. For example, a recombinant human lactoferrin
     (rhLF) gel formulation containing 0.1% to 8.5% rhLF comprised phosphate buffer
    with rhLF 86.67%, Carbopol 980 1.0%, disodium edetate 0.1%, phenoxyethanol
    1.0%, glycerin 4.0%, propylene glycol 5.0%, dimethicone 0.4%, citric acid
    0.0956%, 20% NaOH as needed to pH 6.5-7.5, and water to 100%. RhLF gels
    ranging from 0.1% to 8.5% mediated an improvement in the incidence of 75%
    wound closure of 77% in normal mice at day 12 (p<0.01) and of 66% in
    diabetic db/db mice at day 15 (p<0.05).
ST
    lactoferrin oral topical parenteral wound healing
IT
    Injury
        (bone; oral, parenteral and topical lactoferrin compns. for
       wound treatment)
IT
    Medical goods
       (dressings; oral, parenteral and topical lactoferrin compns.
       for wound treatment)
IT
    Glycosaminoglycans, biological studies
    Polysaccharides, biological studies
    Proteins
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
       (gel containing; oral, parenteral and topical lactoferrin compns.
       for wound treatment)
```

```
IT
     Drug delivery systems
        (gels; oral, parenteral and topical
        lactoferrin compns. for wound treatment)
IT
     Wound
        (infection; oral, parenteral and topical lactoferrin compns.
        for wound treatment)
IT
     Bone, disease
        (injury; oral, parenteral and topical lactoferrin compns. for
        wound treatment)
TΤ
     Antibacterial agents
     Burn
     Drug bioavailability
     Human
     Immunostimulants
     Ulcer
     Wound
     Wound healing promoters
        (oral, parenteral and topical lactoferrin compns. for wound
        treatment)
TΤ
     Chemokines
     Cytokines
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (oral, parenteral and topical lactoferrin compns. for wound
        treatment)
TT
     Lactoferrins
     RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (oral, parenteral and topical lactoferrin compns. for wound
        treatment)
ΙT
     Drug delivery systems
        (oral; oral, parenteral and topical
        lactoferrin compns. for wound treatment)
IT
     Drug delivery systems
        (parenterals; oral, parenteral and topical
        lactoferrin compns. for wound treatment)
IT
     Drug delivery systems
        (topical; oral, parenteral and topical
        lactoferrin compns. for wound treatment)
ΙT
     Diabetes mellitus
        (ulcer from; oral, parenteral and topical lactoferrin compns.
        for wound treatment)
ΙT
     Vein, disease
        (venous stasis ulcer; oral, parenteral and topical lactoferrin
        compns. for wound treatment)
IT
     Digestive tract
     Infection
     Mouth, disease
        (wound; oral, parenteral and topical lactoferrin compns. for
        wound treatment)
ΙT
     120225-54-9, CGS-21680
                              165101-51-9, Regranex
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (combination with; oral, parenteral and topical lactoferrin
        compns. for wound treatment)
                                    9003-05-8, Acrylamide polymer
     9002-89-5, Polyvinyl alcohol
                                                                     9003-39-8,
IT
     Polyvinyl pyrrolidone 106392-12-5, Pluronic 138757-67-2, Carbopol 980
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (gel containing; oral, parenteral and topical lactoferrin compns.
        for wound treatment)
              THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 4
RE
(1) Legrand; Biochem J 1997, V327, P841 HCAPLUS
(2) Nuijens; US 6333311 B1 2001 HCAPLUS
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(3) Valenti; US 5834424 A 1998 HCAPLUS

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(4) van Berkel; Biochemical J 1997, V328, P145 HCAPLUS
    ANSWER 8 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    2003:950872 HCAPLUS
DN
    140:13033
    Entered STN: 07 Dec 2003
ED
ΤI
    Lactoferrin in the treatment of malignant neoplasms and other
    hyperproliferative diseases
    Varadhachary, Atul; Barsky, Rick; Pericle, Frederica; Petrak,
IN
    Karel; Wang, Yenyun
PA
    Agennix Incorporated, USA
    PCT Int. Appl., 51 pp.
SO
    CODEN: PIXXD2
DT
    Patent
    English
T.A
IC
    ICM A61K038-40
    ICS A23L001-305
CC
    1-6 (Pharmacology)
    Section cross-reference(s): 63
FAN.CNT 2
    PATENT NO.
                                        APPLICATION NO.
                      KIND DATE
                                                              DATE
    _____
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                              20031204 WO 2003-US14789 . 20030509 <--
    WO 2003099323
                       A1
PΙ
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
            PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
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            FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                              20040115 US 2003-434769 20030509 <--
    US 2004009895
                        A1
                                       US 2003-435319
    US 2004082504
                              20040429
                                                              20030509 <--
                        Α1
                                        EP 2003-755357
    EP 1507554
                        A1
                              20050223
                                                              20030509 <--
           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
PRAI US 2002-379441P P
                              20020510 <--
    US 2002-379442P
                        Ρ
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    US 2002-379474P
                       P
                              20020510 <--
    WO 2003-US14789
                              20030509
                       W
CLASS
 PATENT NO. CLASS PATENT FAMILY CLASSIFICATION CODES
 _____
WO 2003099323 ICM
                      A61K038-40
                ICS
                      A23L001-305
    The present invention relates to methods of treating a hyperproliferative
AΒ
    disease by administering a composition of lactoferrin alone or in
    combination with standard anti-cancer therapies.
ST
    antitumor lactoferrin neoplasm hyperproliferation disease
    therapy; cancer antitumor human lactoferrin hyperproliferation
    disease therapy
IT
    CD3 (antigen)
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (-pos. T cell; lactoferrin in treatment of malignant
       neoplasms and other hyperproliferative diseases)
IT
    Leukemia
       (acute lymphocytic; lactoferrin in treatment of malignant
       neoplasms and other hyperproliferative diseases)
IT
    Leukemia
       (acute myelogenous; lactoferrin in treatment of malignant
       neoplasms and other hyperproliferative diseases)
```

IT Radiotherapy (and biotherapy; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Antiarteriosclerotics (antiatherosclerotics; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Neuroglia, neoplasm (astrocytoma; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Uterus, neoplasm (cervix; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Leukemia (chronic lymphocytic; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Leukemia (chronic myelomonocytic leukemia; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) TT Intestine, neoplasm (colon; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Neoplasm (fibroma; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Neuroglia, neoplasm (glioblastoma; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Mouth, disease (hairy leukoplakia; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Blood vessel, neoplasm (hemangioma; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Neoplasm (hematopoietic; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Carcinoma (hepatocellular; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Liver, neoplasm (hepatoma; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Intestine, disease (inflammatory; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) IT Drug delivery systems (injections, i.v.; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Adenoma IT Antacids Antiarthritics Antigen-presenting cell Antirheumatic agents Antitumor agents Atherosclerosis Bladder, neoplasm Bone, neoplasm Bos taurus Brain, neoplasm CD4-positive T cell CD8-positive T cell

Carcinoma

Chemotherapy Dendrite (neuron) Digestive tract, neoplasm Head, neoplasm Human Immunostimulants Immunotherapy Kidney, neoplasm Leukemia Lung, neoplasm Lymphoma Mammary gland, neoplasm Melanoma Multiple myeloma Myelodysplastic syndromes Neoplasm Osteoarthritis Ovary, neoplasm Pancreas, neoplasm Prostate gland, neoplasm Psoriasis Rheumatoid arthritis Sarcoma Surgery T cell (lymphocyte) Testis, neoplasm Tongue, neoplasm (lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Interleukin 18 RL: BSU (Biological study, unclassified); BIOL (Biological study) (lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Lactoferrins RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Myoma (leiomyoma; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Adipose tissue, neoplasm (lipoma; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Carcinoma Mesothelium, neoplasm (mesothelioma; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Immune system (mucosal; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Leukemia (myelomonocytic, juvenile; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Lymphocyte (natural killer cell; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Neoplasm (neck; lactoferrin in treatment of malignant neoplasms and other hyperproliferative diseases) Astrocyte (neoplasm, astrocytoma; lactoferrin in treatment of malignant

neoplasms and other hyperproliferative diseases)

IT

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ΙT
     Gingiva
     Hematopoietic precursor cell
     Neck, anatomical
        (neoplasm; lactoferrin in treatment of malignant neoplasms
        and other hyperproliferative diseases)
IT
     Nerve, neoplasm
        (neuroblastoma; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
IT
     Lung, neoplasm
        (non-small-cell carcinoma; lactoferrin in treatment of
        malignant neoplasms and other hyperproliferative diseases)
IT
     Blood vessel, disease
        (occlusion; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
IT
     Drug delivery systems
        (oral; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
IT
     Liver, neoplasm
        (preneoplastic nodule; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
IT
     Carcinoma
        (pulmonary non-small-cell; lactoferrin in treatment of
        malignant neoplasms and other hyperproliferative diseases)
     Carcinoma
IT
        (pulmonary small-cell; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
IT
     Artery, disease
        (restenosis; lactoferrin in treatment of malignant neoplasms
        and other hyperproliferative diseases)
IT
     Eye, neoplasm
        (retinoblastoma; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
IT
     Lung, neoplasm
        (small-cell carcinoma; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
     Carcinoma
IT
        (squamous cell; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
IT
     Drug delivery systems
        (topical; lactoferrin in treatment of malignant
        neoplasms and other hyperproliferative diseases)
IT
     83869-56-1, Granulocyte-macrophage colony-stimulating factor
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (lactoferrin in treatment of malignant neoplasms and other
        hyperproliferative diseases)
ΙT
     15663-27-1, Cisplatin
                             114977-28-5, Docetaxel
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lactoferrin in treatment of malignant neoplasms and other
        hyperproliferative diseases)
              THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Kruzel; US 20030096736 A1 2003 HCAPLUS
(2) Satoh; EP 0730868 A1 1996 HCAPLUS
L92 ANSWER 9 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
     2003:950782 HCAPLUS
AN
DN
     140:752
     Entered STN: 07 Dec 2003
ED
     Oral lactoferrin in the treatment of respiratory disorders
TI
IN
     Glynn, Peter; Varadhachary, Atul
PA
     Agennix Incorporated, USA
     PCT Int. Appl., 56 pp.
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SO

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CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM A61K
CC
     1-9 (Pharmacology)
     Section cross-reference(s): 2, 14, 15, 63
FAN.CNT 1
                                         APPLICATION NO.
                                                             DATE
    PATENT NO.
                      KIND DATE
                               -----
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    WO 2003099207
                        A2
                               20031204 WO 2003-US15763 20030520 <--
PΙ
                               20040408
     WO 2003099207
                        A3
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            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
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            TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
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                               20040115 US 2003-441329
     US 2004009896
                                                                20030520 <--
PRAI US 2002-383280P
                        P
                               20020524 <--
                               20020913 <--
    US 2002-410645P
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CLASS
PATENT NO.
              CLASS PATENT FAMILY CLASSIFICATION CODES
WO 2003099207 ICM
                      A61K
    The invention relates to methods of treating an allergic or non-allergic
     respiratory disorder by administering orally a composition of
     lactoferrin alone or in combination with metal chelators to treat
     respiratory disorders.
ST
     lactoferrin metal chelator interaction allergic nonallergic
     respiratory disorder pharmaceutical; surgery lactoferrin metal
     chelator Tcell respiratory disorder asthma antiasthmatic
    Proteins
IT
    RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (C-reactive, serum; oral lactoferrin for
       treatment of respiratory disorders)
IT
    Antihistamines
        (H1; oral lactoferrin for treatment of respiratory disorders)
IT
    Antibodies and Immunoglobulins
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (IgE, binding inhibitors; oral lactoferrin for treatment of
       respiratory disorders)
IT
    Drug delivery systems
        (aerosols, inhalants, hot air; oral
       lactoferrin for treatment of respiratory disorders)
IT
     Allergy
        (allergic asthma; oral lactoferrin for treatment of
        respiratory disorders)
IT
    Allergy
     Inflammation
    Nose, disease
        (allergic rhinitis; oral lactoferrin for treatment of
       respiratory disorders)
IT
    Asthma
        (allergic; oral lactoferrin for treatment of respiratory
       disorders)
    Antibodies and Immunoglobulins
IT
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (anti-CD23; oral lactoferrin for treatment of respiratory
```

disorders) Bronchi, disease IT Inflammation (bronchitis; oral lactoferrin for treatment of respiratory disorders) IT Drug delivery systems (capsules, enteric-coated; oral lactoferrin for treatment of respiratory disorders) TT Drug delivery systems (carriers; oral lactoferrin for treatment of respiratory disorders) IT Lung, disease (chronic obstructive; oral lactoferrin for treatment of respiratory disorders) Enzymes, biological studies IT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (cleaners; oral lactoferrin for treatment of respiratory disorders) TΤ Mast cell (degranulation agents; oral lactoferrin for treatment of respiratory disorders) IT Allergy (delayed hypersensitivity, associated with atopic or non-atopic asthma; oral lactoferrin for treatment of respiratory disorders) IT Antigen-presenting cell (dendritic; oral lactoferrin for treatment of respiratory disorders) Drug delivery systems IT (enteric; oral lactoferrin for treatment of respiratory disorders) Respiratory tract, disease IT (hyperresponsiveness; oral lactoferrin for treatment of respiratory disorders) IT Drug delivery systems (ligs.; oral lactoferrin for treatment of respiratory disorders) IT Apparatus (mech. breathing device; oral lactoferrin for treatment of respiratory disorders) Atomizing (spraying) TT (moisturized by; oral lactoferrin for treatment of respiratory disorders) Cosmetics IT (moisturizers; oral lactoferrin for treatment of respiratory disorders) IT Drug delivery systems (nasal sprays, salt-water nasal washes or; oral lactoferrin for treatment of respiratory disorders) IT Lymphocyte (natural killer cell; oral lactoferrin for treatment of respiratory disorders) IT Diffusion (of inflammatory cells into lung; oral lactoferrin for treatment of respiratory disorders) IT Allergy Antacids Anti-inflammatory agents Antiasthmatics Antibiotics Antihistamines Antitussives

Asthma

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Blood plasma
Blood serum
Bos taurus
Bronchodilators
CD4-positive T cell
CD8-positive T cell
Chelating agents
Decongestants
Digestive tract
Drug interactions
Emphysema
Eosinophil
Expectorants
Fungicides
Human
Immune system
Immunity
Inflammation
Leukotriene antagonists
Mucous membrane
Respiratory tract, disease
Surgery
T cell (lymphocyte)
   (oral lactoferrin for treatment of respiratory disorders)
CD3 (antigen)
Interleukin 1
Interleukin 10
Interleukin 12
Interleukin 18
Interleukin 2
Interleukin 4
Interleukin 5
VIP receptors
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (oral lactoferrin for treatment of respiratory disorders)
Lactoferrins
RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU
(Therapeutic use); BIOL (Biological study); USES (Uses)
   (oral lactoferrin for treatment of respiratory disorders)
Corticosteroids, biological studies
Glucocorticoids
Steroids, biological studies
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
(Biological study); USES (Uses)
   (oral lactoferrin for treatment of respiratory disorders)
Drug delivery systems
   (oral; oral lactoferrin for treatment of
   respiratory disorders)
Cytokines
RL: BSU (Biological study, unclassified); BIOL (Biological study)
   (pro-inflammatory; oral lactoferrin for treatment
   of respiratory disorders)
Inflammation
Respiratory tract, disease
   (sinusitis, chronic or acute; oral lactoferrin for treatment
   of respiratory disorders)
Drug delivery systems
   (solids; oral lactoferrin for treatment
   of respiratory disorders)
Drug delivery systems
   (sprays, antihistamine; oral lactoferrin
   for treatment of respiratory disorders)
Drug delivery systems
```

TΤ

IT

IT

IT

IT

IT

IT

IT

IT

```
(topical, moisturizing agents; oral
        lactoferrin for treatment of respiratory disorders)
IT
     Adrenoceptor antagonists
        (\alpha-; oral lactoferrin for treatment of respiratory
        disorders)
IT
     Adrenoceptors
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (\alpha-; oral lactoferrin for treatment of respiratory
        disorders)
ΙT
     Adrenoceptor antagonists
        (\beta-; oral lactoferrin for treatment of respiratory
        disorders)
IT
     Interferons
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (\gamma; oral  lactoferrin for treatment of respiratory
        disorders)
IT
     97501-93-4, Tryptase
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (inhibitor; oral lactoferrin for treatment of respiratory
        disorders)
                10102-43-9, Nitric oxide, biological studies
IT
     9041-92-3
                                                                83869-56-1,
     GMCSF
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (oral lactoferrin for treatment of respiratory disorders)
IT
     55-92-5, Methacholine 60-00-4, Ethylenediaminetetraacetic acid,
     biological studies 67-42-5, EGTA 93-14-1, Guaifenesin 150-39-0,
            7681-11-0, Potassium iodide, biological studies
                                                             7782-44-7,
     Oxygen, biological studies
                                85233-19-8, BAPTA
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (oral lactoferrin for treatment of respiratory disorders)
L92
    ANSWER 10 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
     2003:913030 HCAPLUS
DN
     139:358755
ED
     Entered STN: 21 Nov 2003
ΤI
     Intratumorally administered lactoferrin in the treatment of
     malignant neoplasms and other hyperproliferative diseases
IN
     Varadhachary, Atul; Petrak, Karel; Barsky, Rick; O'Malley, Bert
PA
     Agennix Incorporated, USA
     PCT Int. Appl., 45 pp.
so
     CODEN: PIXXD2
DT
     Patent
LA
     English
IC
     ICM A61K038-40
     ICS A23L001-305
     1-6 (Pharmacology)
CC
FAN.CNT 2
                                          APPLICATION NO.
     PATENT NO.
                        KIND
                               DATE
                                                                   DATE
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     WO 2003094952
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                                          WO 2003-US14584
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            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
            PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
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         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     US 2004009895
                                20040115
                                         US 2003-434769
                         A1
                                                                  20030509 <--
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20040429

Α1

US 2003-435319

20030509 <--

US 2004082504

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PRAI US 2002-379441P
                        P
                               20020510 <--
                        P
    US 2002-379442P
                               20020510 <--
    US 2002-379474P
                        P
                               20020510 <--
CLASS
               CLASS PATENT FAMILY CLASSIFICATION CODES
PATENT NO.
 ______
WO 2003094952 ICM
                      A61K038-40
                ICS
                      A23L001-305
    The present invention relates to methods of treating a hyperproliferative
AB
    disease by administering a composition of lactoferrin alone or in
    combination with standard anticancer therapies.
ST
    intratumor lactoferrin cancer hyperproliferative disease
ΙT
    Antiarteriosclerotics
        (antiatherosclerotics; intratumorally administered
       lactoferrin in the treatment of malignant neoplasms and other
       hyperproliferative diseases)
    Neuroglia, neoplasm
IT
        (astrocytoma; intratumorally administered lactoferrin in the
       treatment of malignant neoplasms and other hyperproliferative diseases)
IT
    Gingiva
        (cancer; intratumorally administered lactoferrin in the
       treatment of malignant neoplasms and other hyperproliferative diseases)
IT
    Uterus, neoplasm
        (cervix; intratumorally administered lactoferrin in the
       treatment of malignant neoplasms and other hyperproliferative diseases)
IT
    Intestine, neoplasm
        (colon; intratumorally administered lactoferrin in the
       treatment of malignant neoplasms and other hyperproliferative diseases)
TT
    Neoplasm
        (fibroma; intratumorally administered lactoferrin in the
       treatment of malignant neoplasms and other hyperproliferative diseases)
    Neuroglia, neoplasm
IT
        (glioblastoma; intratumorally administered lactoferrin in the
       treatment of malignant neoplasms and other hyperproliferative diseases)
IT
    Blood vessel, neoplasm
        (hemangioma; intratumorally administered lactoferrin
       in the treatment of malignant neoplasms and other hyperproliferative
       diseases)
IT'
    Intestine, disease
        (inflammatory; intratumorally administered lactoferrin in the
       treatment of malignant neoplasms and other hyperproliferative diseases)
TT
    Adenoma
    Antirheumatic agents
    Antitumor agents
      Atherosclerosis
    Bladder, neoplasm
    Bone, neoplasm
    Brain, neoplasm
    Digestive tract, neoplasm
    Head, neoplasm
    Human
    Immunotherapy
    Kidney, neoplasm
    Leukemia
    Lung, neoplasm
    Lymphoma
    Mammary gland, neoplasm
    Osteoarthritis
    Ovary, neoplasm
    Pancreas, neoplasm
    Prostate gland, neoplasm
    Psoriasis
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Radiotherapy

Rheumatoid arthritis Sarcoma Surgery T cell (lymphocyte) Testis, neoplasm Tongue, neoplasm (intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) TΤ Interleukin 18 RL: BSU (Biological study, unclassified); BIOL (Biological study) (intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Lactoferrins RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Myoma (leiomyoma; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) Mouth, disease TΤ (leukoplakia; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Adipose tissue, neoplasm (lipoma; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Carcinoma Mesothelium, neoplasm (mesothelioma; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) TΤ Lymphocyte (natural killer cell; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Neoplasm (neck; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Astrocyte (neoplasm, astrocytoma; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) Neck, anatomical IT (neoplasm; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Nerve, neoplasm (neuroblastoma; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Blood vessel, disease (occlusion; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) TΤ Antiarthritics (osteoarthritis; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) IT Artery, disease (restenosis; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative diseases) Eye, neoplasm IT (retinoblastoma; intratumorally administered lactoferrin in the treatment of malignant neoplasms and other hyperproliferative

diseases)

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IT
    83869-56-1, Granulocyte-macrophage colony-stimulating factor
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (intratumorally administered lactoferrin in the treatment of
       malignant neoplasms and other hyperproliferative diseases)
             THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Kruzel; US 20030096736 A1 2003 HCAPLUS
(2) Satoh; EP 0730868 A1 1996 HCAPLUS
L92
    ANSWER 11 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
AN
    2003:696760 HCAPLUS
DN
    139:219356
ED
    Entered STN: 05 Sep 2003
ΤI
    Pharmaceutical composition for treatment of vascular disease or states of
    tissue hypoperfusion with hypoxic and/or ischemic consequences
IN
    Norrby, Klas
PA
    Swed.
    PCT Int. Appl., 27 pp.
SO
    CODEN: PIXXD2
DT
    Patent
    English
LA
    ICM A61K038-40
IC
    ICS A61P009-00
    63-6 (Pharmaceuticals)
CC
    Section cross-reference(s): 1
FAN.CNT 1
    PATENT NO.
                                        APPLICATION NO.
                      KIND DATE
                                                               DATE
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PΙ
    WO 2003072129
                        A1
                              20030904 WO 2003-SE329
                                                               20030227 <--
        W: AU, JP, US
        RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
            IT, LU, MC, NL, PT, SE, SI, SK, TR
    EP 1478387
                        A1
                              20041124 EP 2003-743090
                                                                20030227 <--
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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CLASS
 PATENT NO.
              CLASS PATENT FAMILY CLASSIFICATION CODES
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 WO 2003072129 ICM
                       A61K038-40
                TCS
                      A61P009-00
AB
    Disclosed is the use of a substance selected from the group consisting of
    human apolactoferrin and/or peptides derivable from human
    lactoferrin and/or natural metabolites of human
    lactoferrin and/or functionally equivalent analogs of human
    apolactoferrin for the production of a pharmaceutical composition for treatment
    and/or prevention of a vascular disease and/or states of tissue
    hypoperfusion with hypoxic and/or ischemic consequences. Thus, oral or
    s.c. administration of apolactoferrin specifically enhanced the
    VEGF-mediated angiogenesis.
ST
    apolactoferrin pharmaceutical vascular disease; tissue hypoperfusion
    hypoxia apolactoferrin; ischemia apolactoferrin pharmaceutical
    Heart, disease
IT
        (angina pectoris; pharmaceutical compns. for
       treatment of vascular disease or states of tissue hypoperfusion with
       hypoxic and/or ischemic consequences)
IT
    Lactoferrins
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (apolactoferrins; pharmaceutical compns. for treatment of
       vascular disease or states of tissue hypoperfusion with hypoxic and/or
```

ischemic consequences)

IT Ischemia

(cardiac; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Necrosis

(gangrene; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Heart, disease

(infarction; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Drug delivery systems

(inhalants; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Heart, disease

(ischemia; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Ulcer

(leg; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Drug delivery systems

(oral; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Drug delivery systems

(parenterals; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Ulcer

(peptic; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Artery, disease

(peripheral, occlusion; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Alopecia

Angiogenesis

Blood vessel, disease

Human

Hypoxia, animal

Perfusion

(pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Lactoferrins

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Drug delivery systems

(topical; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Digestive tract, disease

(ulcer, peptic; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences)

IT Leg, disease

(ulcer; pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences) IT 127464-60-2, Vascular endothelial growth factor RL: ADV (Adverse effect, including toxicity); BIOL (Biological study) (pharmaceutical compns. for treatment of vascular disease or states of tissue hypoperfusion with hypoxic and/or ischemic consequences) RE.CNT THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD (1) Alpharma As; WO 0012541 A2 2000 HCAPLUS (2) Anon; PATENT ABSTRACTS OF JAPAN 1996, V199 (602) (3) Biocem Sa; JP 2001527406 TT 2001 HCAPLUS (4) Morinaga Milk Ind Co Ltd; JP 7278011 A 1995 (5) Morinaga Milk Industry Co Ltd; JP 09194388 A2 1997 HCAPLUS (6) Nakajima, M; Journal of cellular physiology 1997, V170(2), PP101 (7) Pharming B V; WO 9833509 A2 1998 HCAPLUS (8) Science Invest Ab; WO 0001730 Al 2000 HCAPLUS (9) Shimura, S; Investigative ophthalmology & visual science (UNITED STATES) 1998, V39(8), Pp1346 ANSWER 12 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN L92 AN2003:551399 HCAPLUS DN 139:90498 ED Entered STN: 18 Jul 2003 TI Compositions for improving lipid metabolism Harada, Etsumori; Takeuchi, Takashi; Ando, Kunio; Shimizu, Hirohiko TN Nuclear Receptor Ligand Co., Ltd., Japan PA so PCT Int. Appl., 51 pp. CODEN: PIXXD2 DT Patent T.A Japanese IC ICM A61K038-40 A61K038-16; A61K009-14; A61K009-16; A61K009-20; A61K009-48; A61P001-16; A61P003-04; A61P003-06; A61P003-10; A61P009-12 CC 63-6 (Pharmaceuticals) Section cross-reference(s): 1, 14 FAN.CNT 1 PATENT NO. KIND DATE APPLICATION NO. DATE _____ -----____ -----PΙ WO 2003057245 A1 20030717 WO 2002-JP13858 20021227 <--W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR; CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG 20041013 EP 2002-793463 EP 1466621 A1 20021227 <--AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK US 2005020484 20050127 US 2004-500245 A1 20040625 <--PRAI JP 2001-400641 20011228 Α <--WO 2002-JP13858 W 20021227 CLASS CLASS PATENT FAMILY CLASSIFICATION CODES PATENT NO. _______ _ _ _ _ WO 2003057245 ICM A61K038-40 A61K038-16; A61K009-14; A61K009-16; A61K009-20; ICS A61K009-48; A61P001-16; A61P003-04; A61P003-06; A61P003-10; A61P009-12

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EP 1466621
                 ECLA
                        A61K038/01D; A61K038/01D6; A61K038/40
                        A61K038/01D; A61K038/01D6; A61K038/40
US 2005020484
                 ECLA
                                                                             <--
    A medicinal composition contains as the active ingredient at least one member
     selected from the group consisting of lactoferrin proteins
     including lactoferrin and conalbumin and enzymically digested
    products of lactoferrin proteins including lactoferricin and
    peptides of conalbumin corresponding to lactoferricin. The composition is
     useful for improving lipid metabolism For example, it is useful in treating
     lifestyle-related diseases such as hypercholesterolemia,
     hypertriglyceridemia, low-d. lipoprotein hypercholesterolemia,
     high-d. lipoprotein hypocholesterolemia, obesity, fat liver,
     cholesterol cholelithiasis, severe obesity, hyperlipidemia,
     hypertension, type II diabetes. The composition can elevate basal metabolic
     rate.
ST
     lipid metab drug disease life style habit
     Lipids, biological studies
IT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (drugs for improving lipid metabolism)
IT
     Calculi, biliary
       Hypercholesterolemia
       Hypertension
       Hypertriglyceridemia
     Obesity
        (drugs for improving lipid metabolism for treatment of)
IT
    Liver
        (fat; drugs for improving lipid metabolism for treatment of)
IT
     Lipids, biological studies
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (hyperlipidemia; drugs for improving lipid metabolism for treatment of)
TΤ
     Lactoferrins
     Peptides, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (in drugs for improving lipid metabolism)
    Disease, animal
IT
        (lifestyle-related; drugs for improving lipid metabolism for treatment of)
TΤ
    Diabetes mellitus
        (non-insulin-dependent; drugs for improving lipid metabolism for treatment
        of)
TΤ
    Liver
        (toxicity, fat; drugs for improving lipid metabolism for treatment of)
     1391-06-6, Conalbumin 146897-68-9, Lactoferricin
TΤ
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (in drugs for improving lipid metabolism)
              THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
       24
RE
(1) Agennix Inc; CN 1262625 A 1998
(2) Agennix Inc; JP 2001519815 A 1998
(3) Agennix Inc; NZ 500712 A 1998 HCAPLUS
(4) Agennix Inc; EP 979099 A1 1998 HCAPLUS
(5) Agennix Inc; WO 9844940 A1 1998 HCAPLUS
(6) Agennix Inc; AU 9869647 A 1998 HCAPLUS
(7) Agennix Inc; MX 9909240 A1 1998 HCAPLUS
(8) Biotech Australia Pty Ltd; WO 0022909 A2 2000 HCAPLUS
(9) Biotech Australia Pty Ltd; AU 200010712 A 2000
(10) Bukh Meditec AS; JP 05-500668 A 1991
(11) Bukh Meditec AS; JP 2927950 B2 1991 HCAPLUS
(12) Bukh Meditec AS; EP 493513 B1 1991 HCAPLUS
(13) Bukh Meditec AS; US 5213808 A 1991 HCAPLUS
(14) Bukh Meditec AS; DE 69009769 E 1991
(15) Bukh Meditec AS; AU 9065051 A 1991 HCAPLUS
(16) Bukh Meditec AS; WO 9104015 A1 1991 HCAPLUS
(17) Meiji Milk Products Co Ltd; JP 2000325046 A 2000 HCAPLUS
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(18) Morinaga Milk Industry Co Ltd; JP 05-176713 A 1993 HCAPLUS

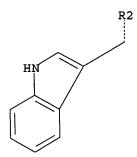
- (19) Morinaga Milk Industry Co Ltd; KR 2000070051 A 1999
- (20) Morinaga Milk Industry Co Ltd; US 6319895 B1 1999 HCAPLUS
- (21) Morinaga Milk Industry Co Ltd; EP 955058 Al 1999 HCAPLUS
- (22) Morinaga Milk Industry Co Ltd; WO 9830235 A1 1999 HCAPLUS
- (23) Morinaga Milk Industry Co Ltd; JP 2001048808 A 2001 HCAPLUS
- (24) Tokiwa Chemical Industries Ltd; JP 2000198739 A 2000 HCAPLUS
- IT 146897-68-9, Lactoferricin
 - RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (in drugs for improving lipid metabolism)
- RN 146897-68-9 HCAPLUS
- CN L-Phenylalanine, L-phenylalanyl-L-lysyl-L-cysteinyl-L-arginyl-L-arginyl-L-tryptophyl-L-arginyl-L-arginyl-L-methionyl-L-lysyl-L-leucylglycyl-L-alanyl-L-prolyl-L-seryl-L-isoleucyl-L-threonyl-L-cysteinyl-L-valyl-L-arginyl-L-arginyl-L-alanyl-, cyclic (3→20)-disulfide (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PAGE 1-A

Me S NH S Photo NH
$$(CH_2)_3$$
 S $(CH_2)_3$ $(CH_2)_3$

PAGE 4-A



L92 ANSWER 13 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2002:613228 HCAPLUS

ED Entered STN: 16 Aug 2002.

TI New ingredients from dairy foods

AU Morgan, Wendy

CS N/A, North Sydney, NSW 2059, Australia

SO Abstracts of Papers, 224th ACS National Meeting, Boston, MA, United States, August 18-22, 2002 (2002), AGFD-041 Publisher: American Chemical Society, Washington, D. C. CODEN: 69CZPZ

DT Conference; Meeting Abstract

LA English

AB Dairy foods have traditionally been considered a source of nutrition, particularly high quality protein, calcium and other minerals, and vitamins. In the last few decades, milk and other dairy foods have been maligned due to their saturated fat and cholesterol contents and the belief that these constituents increase the risk of coronary heart disease. There are several studies, which indicate that dairy products may not potentiate atherosclerosis. In fact there are factors in milk that may actively protect against heart disease such as calcium, bioactive peptides, folic acid, vitamin B6, vitamin B12 and conjugated linoleic acid. A range of other activities has been demonstrated for dairy components. Milk proteins are an important source of bioactive peptides showing opioid and ACE-inhibitory activity. Lactoferrin has been shown to have a bifidus effect and antimicrobial activity. also improves iron bioavailability. Glycomacropeptide, a-lactoglobulin, a-lactalbumin and casein phosphopeptides affect physiol. functions. The development of membrane technologies allows the fractionation of milk proteins to produce a range of products with potential impact on human health.

L92 ANSWER 14 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2000:444357 HCAPLUS

DN 133:250634

ED Entered STN: 04 Jul 2000

TI LDL receptor-related protein mediates uptake of aggregated LDL in human vascular smooth muscle cells

AU Llorente-Cortes, Vicenta; Martinez-Gonzalez, Jose; Badimon, Lina

CS Cardiovascular Research Center, Institut de Recerca de l'Hospital de la Santa Creu i Sant Pau, Barcelona, Spain

SO Arteriosclerosis, Thrombosis, and Vascular Biology (2000), 20(6), 1572-1579

CODEN: ATVBFA; ISSN: 1079-5642

PB Lippincott Williams & Wilkins

DT Journal

LA English

CC 14-5 (Mammalian Pathological Biochemistry)

AB Foam cell formation is a key event in the onset and progression of atherosclerotic lesions. We have previously reported that internalization of aggregated low d. lipoproteins (agLDLs) by vascular smooth muscle cells (VSMCs) produces cholesteryl ester (CE) accumulation in these cells. The aim of this study was to analyze whether the low d. lipoprotein receptor-related protein (LRP) mediates the uptake of agLDL by VSMCs. First, immunocytochem. and fluorescence microscopic anal. with the use of anti-LRP antibodies indicated that there was a high expression of LRP in VSMCs. Confocal microscopic anal. with the use of agLDLs labeled with fluorochrome 1,1'-dioctadecyl-3,3,3',3'tetramethylindocarbocyanine and anti-LRP antibodies showed the colocalization of agLDL and LRP. The second approach was to analyze the effect of LRP ligands on agLDL internalization; lactoferrin strongly inhibited CE accumulation from agLDLs (85.0±5.7% at 25 μg/mL) by impairing agLDL binding. Coincubation of agLDL with anti-LRP antibodies decreased in a dose-dependent manner agLDL-derived CE accumulation (from 20% at 12.5 $\mu g/mL$ to 80% at 50 $\mu g/mL)\,.$ The third approach was to evaluate whether antisense LRP oligodeoxynucleotides were able to block agLDL internalization. Treatment of VSMCs with 5 µmol/L antisense LRP oligodeoxynucleotides reduced agLDL-derived CE accumulation by 84±2%. In conclusion, these results from immunol., biochem., and mol. interventions demonstrate that LRP mediates the binding and internalization of agLDL in human VSMCs. Because LRP is highly expressed in VSMCs and the uptake of 1 LDL aggregate amts. to the deposition of several hundreds of LDL particles, the uptake of agLDL through LRP could have a crucial role for lipid deposition in VSMCs.

ST LRP aggregated LDL uptake cholesterol accumulation vessel muscle atherosclerosis

IT Molecular association

(LDL receptor-related protein mediates binding and internalization of aggregated LDL, and **cholesterol** ester accumulation in human vascular smooth muscle cells)

IT Lipoproteins

RL: BOC (Biological occurrence); BSU (Biological study, unclassified);
BIOL (Biological study); OCCU (Occurrence)
 (low-d.; LDL receptor-related protein

mediates uptake of aggregated **LDL** in human vascular smooth muscle cells)

IT Blood vessel

(smooth muscle; LDL receptor-related protein mediates uptake of aggregated LDL in human vascular smooth muscle cells)

IT Atherosclerosis

(uptake of aggregated LDL through LDL receptor-related protein could have crucial role for lipid deposition in human vascular smooth muscle cells and onset of atherosclerotic lesions)

IT Biological transport

(uptake; LDL receptor-related protein mediates uptake of aggregated LDL in human vascular smooth muscle cells)

IT Receptors

RL: BAC (Biological activity or effector, except adverse); BOC (Biological occurrence); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); OCCU (Occurrence); PROC (Process) (α2-macroglobulin; LDL receptor-related protein mediates uptake of aggregated LDL in human vascular smooth muscle cells)

IT 57-88-5D, Cholesterol, esters

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(LDL receptor-related protein mediates binding and internalization of aggregated LDL, and **cholesterol** ester accumulation in human vascular smooth muscle cells)

RE.CNT 56 THERE ARE 56 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

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- IT 57-88-5D, Cholesterol, esters
 - RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)
 - (LDL receptor-related protein mediates binding and internalization of aggregated LDL, and **cholesterol** ester accumulation in human vascular smooth muscle cells)
- RN 57-88-5 HCAPLUS

CN Cholest-5-en-3-ol (3\beta) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.

PATENT NO.

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ED
     Entered STN: 19 May 2000
TI
     Pharmaceutical preparation containing a receptor antagonist for treating
     blood-clotting disorders
IN
     Schwarz, Hans-Peter; Turecek, Peter; Binder, Bernd
PA
     Baxter Aktiengesellschaft, Austria
SO
     PCT Int. Appl., 20 pp.
     CODEN: PIXXD2
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          A61K038-57; A61K038-49; A61K038-40; A61K038-16
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CLASS
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CLASS PATENT FAMILY CLASSIFICATION CODES

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ICS A61K038-36; A61K038-48; A61P007-04; A61K038-37;
A61K038-17; A61K038-57; A61K038-49; A61K038-40;
A61K038-16
WO 2000027425 ECLA A01K067/027A3A; A61K038/16+M; A61K038/37+M;

A61K038/48K+M; A61K038/57+M

AB A pharmaceutical preparation for treating blood-clotting disorders by increasing the biol. half-life of blood-coaqulation factors contains ≥1 blood-coagulation factor pro-protein and a coagulation factor receptor-binding competitor (ligand) which is inert with regard to its effects on blood clotting. By preventing the internalization and degradation of the coagulation factor via the receptor, the competitor prolongs the functional half-life of the factor in vivo. The pro-protein may be Factor II, V, VII, VIII, IX, X, XI, XII, protein C, or especially von Willebrand factor. For example, the binding of activated Factor VIII to lipoprotein receptor-related protein (LRP), which is responsible for binding, internalization, and degradation of Factor VIII, is inhibited by receptor-associated protein (RAP). Thus, in mice with severe Factor VIII deficiency which were administered recombinant human Factor VIII, administration of a RAP fusion protein with glutathione S-transferase 10 min prior to Factor VIII administration greatly increased the plasma Factor VIII antigen concentration 1 h later.

ST blood coagulation factor degrdn inhibition; receptor assocd protein coagulation factor; coagulopathy treatment receptor ligand

IT Apolipoproteins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(E, **VLDL** enriched with; pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)

IT Glycoproteins, specific or class

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(PAAG (pregnancy-associated $\alpha 2$ -glycoprotein), complexes with proteinase; pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)

IT Proteins, specific or class

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(RAP (receptor-associated protein); pharmaceutical preparation containing receptor

antagonist for treating blood-clotting disorders)

IT Blood coagulation

(disorder; pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)

IT Toxins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(exotoxin A, of Pseudomonas; pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)

IT Ligands

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(for blood-coagulation factor receptors; pharmaceutical preparation containing

receptor antagonist for treating blood-clotting disorders)

IT Receptors

RL: BAC (Biological activity or effector, except adverse); BPR (Biological

process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

(for blood-coagulation factor; pharmaceutical preparation containing receptor

antagonist for treating blood-clotting disorders)

IT Lipoproteins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(low-d.; pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)

IT Proteins, specific or class

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(matrix; pharmaceutical preparation containing receptor antagonist for treating

blood-clotting disorders)

IT Animal virus

Rhinovirus

(pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)

IT Enzymes, biological studies

Lactoferrins

Lipoproteins

Thrombospondins

Toxins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)

IT Blood-coagulation factors

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(precursors; pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)

IT Lipoproteins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(very-low-d. β -; pharmaceutical

preparation containing receptor antagonist for treating blood-clotting disorders)

IT Lipoproteins

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(very-low-d., lipoprotein

lipase-enriched; pharmaceutical preparation containing receptor antagonist

for

treating blood-clotting disorders)

IT Receptors

RL: BAC (Biological activity or effector, except adverse); BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)

 $(\alpha 2\text{-macroglobulin}, \text{ ligands for; pharmaceutical preparation containing receptor antagonist for treating blood-clotting disorders)}$

IT 9001-01-8, Kallikrein 9001-24-5, Blood-coagulation factor V 9001-25-6, Blood-coagulation factor VII 9001-26-7, Blood-coagulation factor II 9001-27-8 9001-28-9, Blood-coagulation factor IX 9001-29-0, Blood-coagulation factor X 9001-30-3, Blood-coagulation factor XII

9001-92-7D, Proteinase, complexes with α 2-macroglobulin

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9002-04-4D, Thrombin, complexes with plasminogen activator inhibitor 1
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    antitrypsin 9013-55-2, Blood-coagulation factor XI 9039-53-6,
    Urokinase 9041-92-3D, α1-Antitrypsin, complexes with elastase
    9087-70-1, Aprotinin 50812-37-8D, Glutathione S-transferase, fusion
    products with receptor-associated protein 60202-16-6, Blood-coagulation
    factor XIV 81604-65-1D, Heparin cofactor II, complexes with thrombin
    82657-92-9, Prourokinase 109319-16-6 139639-23-9, Tissue-type
    plasminogen activator 140208-23-7, Plasminogen activator inhibitor 1
    148196-69-4D, Protease-nexin 1, complexes with urokinase
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       (pharmaceutical preparation containing receptor antagonist for treating
       blood-clotting disorders)
L92 ANSWER 16 OF 16 HCAPLUS COPYRIGHT 2005 ACS on STN
    1994:708429 HCAPLUS
    121:308429
    Entered STN: 24 Dec 1994
    Modified low-density lipoprotein-(LDL) binding agents
    Matsuda, Ichiro; Oota, Takao; Tomita, Mamoru; Shimamura, Seiichi; Kawase,
    Kozo; Takase, Mitsunori; Kajikawa, Mikio
    Morinaga Milk Industry Co Ltd, Japan
    Jpn. Kokai Tokkyo Koho, 8 pp.
    CODEN: JKXXAF
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    ICM A61K037-14
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    63-8 (Pharmaceuticals)
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FAN.CNT 1
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                      A61K037-18; A61M001-36
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    Lactoferrin and/or its hydrolyzates are useful as binding agent
    to modified LDL for treatment of arteriosclerosis. Bovine
    lactoferrin was treated with acetylated human LDL to show good
    binding activity. Lactoferrin-containing tablets and freezed-dried
    preparation and immobilized lactoferrin for hemodialysis were manufactured
    antiarteriosclerotic LDL binder lactoferrin hemodialysis
    Antiarteriosclerotics
        (lactoferrin (hydrolyzates) as binder for modified LDL for
       treatment of arteriosclerosis)
    Lactoferrins
    RL: BPR (Biological process); BSU (Biological study, unclassified); THU
     (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
        (lactoferrin (hydrolyzates) as binder for modified
       LDL for treatment of arteriosclerosis)
    Dialysis
        (hemo-, lactoferrin (hydrolyzates) as binder for modified LDL
       for treatment of arteriosclerosis)
    Lactoferrins
```

RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (hydrolyzates, lactoferrin (hydrolyzates)

as hinder for modified LDL for treatment of arteriosclerosis)

as binder for modified LDL for treatment of arteriosclerosis)

IT Lipoproteins

RL: BPR (Biological process); BSU (Biological study, unclassified); REM
(Removal or disposal); BIOL (Biological study); PROC (Process)
 (low-d., acetylated, lactoferrin
 (hydrolyzates) as binder for modified LDL for treatment of
 arteriosclerosis)

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- L116 ANSWER 1 OF 7 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

AN 2004-460986 [43] WPIX

DNC C2004-172138

- TI Treating a cardiovascular disease comprises administering to a subject an effective amount of a **lactoferrin** composition to provide an improvement in the cardiovascular disease in the subject.
- DC B04 D16
- IN ENGELMAYER, J; GLYNN, P; VARADHACHARY, A; WANG, Y
- PA (ENGE-I) ENGELMAYER J; (GLYN-I) GLYNN P; (VARA-I) VARADHACHARY A; (WANG-I) WANG Y; (AGEN-N) AGENNIX INC

CYC 107

PI WO 2004050037 A2 20040617 (200443)* EN 38 A61K000-00 RW: AT BE BG BW CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE

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AB
     NOVELTY - Treating a cardiovascular disease comprises administering to a
     subject an effective amount of a lactoferrin composition to
     provide an improvement in the cardiovascular disease in the subject.
          DETAILED DESCRIPTION - An INDEPENDENT CLAIM is included for a method
     of modulating atherosclerosis in a subject comprising administering to the
     subject an effective amount of a lactoferrin composition to
     modulate atherosclerosis in the subject.
          ACTIVITY - Cardiant; Antiarteriosclerotic. No biological data given.
          MECHANISM OF ACTION - Gene therapy; HMG-coA reductase inhibitor.
          USE - The method is useful for treating a cardiovascular disease,
     e.g. atherosclerosis (claimed).
     Dwg.0/5
FS
     CPI
    AB; DCN
FA
MC
     CPI: B04-N02; B04-N0200E; B04-N06; B04-N0600E; B06-D01; B07-A02B; B07-A03;
          B07-D02; B07-D04C; B10-A22; B10-B01B; B10-C03; B10-C04A; B10-F02;
          B14-C03; B14-D02A2; B14-D05D; B14-F01;
          B14-F02; B14-F06; B14-F07; B14-S03;
          D05-C12; D05-H17A6
TECH
                    UPTX: 20040709
     TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Method: In treating a
     cardiovascular disease, the cardiovascular disease is atherosclerosis. The
     lactoferrin composition reduces levels of circulating total
     cholesterol, low-density lipoproteins (LDL), very low-density lipoproteins
     (VLDL), or triglycerides in the subject. The lactoferrin
     composition increases the levels of circulating high-density lipoproteins
     (HDL) in the subject. The lactoferrin composition reduces the
     levels of vascular inflammation, circulating C-reactive protein (CRP),
     proliferation of vascular smooth muscle cells, vascular spasm or vascular
     hyper-reactivity in the subject. The lactoferrin composition
     promotes endothelial integrity or healing in the subject. The
     lactoferrin composition is dispersed in a carrier. The
     lactoferrin is mammalian lactoferrin. The
     lactoferrin is human or bovine. The lactoferrin is
     recombinant lactoferrin. The lactoferrin composition
     comprises an N-terminal lactoferrin variant. The N-terminal
     lactoferrin variant lacks at least the N-terminal glycine residue.
     The N-terminal lactoferrin variant comprises at least 1% to at
     least 50% of the lactoferrin composition. The
     lactoferrin composition reduces the production or activity of
     pro-inflammatory cytokines. The method further comprises administering a
     lactoferrin composition in combination with an anti-cholesterol
     agent or an anti-inflammatory agent. The anti-cholesterol agent is
     selected from cholesterol absorption inhibitors, bile acid sequestrants,
     nicotinic acid, fibric acids and HMG-coA reductase inhibitors. The bile
     acid sequestrants are selected from cholestyramine, colestipol and
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colesevalam. The fibric acids are selected from gemfibrozil, fenofibrate and clofibrate. The HMG-coA reductase inhibitors are selected from lovastatin, pravastatin, simvastatin, fluvastatin, atorvastatin and cerivastatin. In modulating atherosclerosis in a subject, the modulating is reducing the incidence or severity of atherosclerosis in the subject. ABEX UPTX: 20040709 ADMINISTRATION - Dosage is 1 ng-20 q per day or 0.1-5 q per day. The lactoferrin composition is administered parenterally, e.g. subcutaneously, intramuscularly, intraperitoneally, intravenously, intraarterially, intramyocardially, transendocardially, transepicardially, or intrathecally, or orally (all claimed). L116 ANSWER 2 OF 7 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN 2004-247885 [23] WPIX C2004-096768 New peptidyl analogs are growth hormone secretagogue agonists useful in the treatment of e.g. weight loss, sexual dysfunction, diabetes or cardiovascular diseases. B04 C03 ZHENG, X D (SCRC) SCRAS SOC CONSEILS RECH & APPL SCI CYC WO 2004014415 A1 20040219 (200423)* EN 77 A61K038-40 RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PG PH PL PT RO RU SC SD SE SG SK SL SY TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW AU 2003259062 A1 20040225 (200456) A61K038-40 WO 2004014415 A1 WO 2003-US24834 20030808; AU 2003259062 A1 AU 2003-259062 20030808 AU 2003259062 A1 Based on WO 2004014415 FDT PRAI US 2002-402263P 20020809 ICM A61K038-40 WO2004014415 A UPAB: 20040405 NOVELTY - Peptidyl analogs (I) or their salts are new. DETAILED DESCRIPTION - Peptidyl analogs of formula R1- A1- A2- A3-A4- A5-R2 (I) or their salts are new. A1 = Aib, amino piperidinylcarboxylic acid (Apc) or isonipecotic acid (Inp); A2 and A3 = S1, beta -(1-naphthyl)-D-alanine (D-1Nal), beta -(2-naphthyl)-D-alanine (D-2Nal), D-Ser(Bzl) or D-Trp; S1 = 3-benzoylthienyl-D-alanine (D-Bal), 4,4'-biphenyl-D-alanine (D-Bip), 4-benzoyl-D-phenylalanine (D-Bpa) or beta, beta -diphenyl-D-alanine (D-Dip); A4 = S2, S4, Phe or 2'-(4-phenyl)imidazolyl (Pim); S2 = beta -(2-furyl)-alanine (2Fua), pentafluorophenylalanine (Pff), beta - (4-thiazolyl) alanine (Taz), or Thr(Bzl); S4 = beta -(2-thienyl)alanine (2Thi), beta -(3-thienyl)alanine (3Thi), Orn, beta -(2-pyridyl)alanine (2Pal), beta -(3-pyridyl)alanine (3Pal) or beta - (4-pyridyl) alanine (4Pal); A5 = Apc, S3 or none;S3 = 2,4-diaminobutyric acid (Dab), 2,3-diaminopropionic acid (Dap), Lys or Orn; R1 = H, 1-6C alkyl, 5-14C aryl, 1-6C alkyl-5-14C aryl, 3-8C cycloalkyl or 2-10C acyl; and R2 = OH, NH2.

DC

IN

PA

ADT

IC

AB

Provided that:

(a) when A5 is S3, then A2 or A3 is S1 or A4 is S2; and (b) when A5 is absent then A3 is D-Bip, D-Bpa or D-Dip or A4 is S2 or A1 is Apc and A2 or A3 is S1 or A4 is S4.

ACTIVITY - Antidiabetic; Anorectic; Cardiovascular-Gen.; Cardiant; Osteopathic; Endocrine-Gen.; Cytostatic; Immunomodulator; Muscular-Gen.; Opthalmological.

MECHANISM OF ACTION - Growth hormone secretagogue (GHS) agonist; Ghrelin agonist; Ghrelin antagonist; Growth hormone secretion stimulator; Growth hormone secretion antagonist.

Test details are described but no results are given. USE - Used:

- (1) in the treatment of a growth hormone deficient state, for facilitating weight gain, maintenance of weight and/or appetite increase in a disease or disorder accompanied by weight loss (including anorexia, bulimia, cancer cachexia, AIDS, AIDS wasting, cachexia, wasting in frail elderly, chemotherapy, radiation therapy, immobilization or dialysis);
- (2) for increasing muscle mass and bone density, sexual dysfunction, maintenance of weight for physical functioning, recovery of physical function;
- (3) for facilitating weight loss and treating obesity due to a disease or condition (including hypertension, diabetes, dyslipidemia, cardiovascular disease, gall stones, osteoarthritis or cancers);
 - (4) for appetite decrease, weight maintenance;
- (5) for diabetes, complications of diabetes including retinopathy, cardiovascular disorders;
- (6) for achieving a beneficial cardiovascular effect (e.g. inhibition of apoptosis of cardiomyocytes, cardiac endothelial cells or vascular endothelial cells, attenuation of the development of cardiac cachexia, reduction in systemic vascular resistance, an increase in cardiac output or improvement of cardiac structure or function) in human;
- (7) for chronic/severe heart failure and for eliciting ghrelin agonist/antagonist effect (claimed).

Also useful for facilitating weight gain in farm animals (e.g. pigs, cows or chickens).

ADVANTAGE - The compounds are potent growth hormone secretagogues. $\ensuremath{\text{Dwg.0/0}}$

FS CPI

FA AB; GI; DCN

MC CPI: B04-C01A; B14-D01; B14-E11; B14-E12; B14-F01;

B14-F02; B14-L01; B14-L06; B14-N01; B14-P02; B14-S04;

C04-C01A; C14-D01; C14-E11; C14-E12; C14-F01;

C14-F02; C14-L01; C14-L06; C14-N01; C14-P02; C14-S04

TECH

TECHNOLOGY FOCUS - ORGANIC CHEMISTRY - Preparation: (I) Is prepared by using standard solid phase peptide synthesis reported in Stewart, J.M. et al., Solid Phase Synthesis (Pierce Chemical Co., 2nd ed. 1984).

ABEX UPTX: 20040405

SPECIFIC COMPOUNDS - 180 Compounds are specifical

UPTX: 20040405

SPECIFIC COMPOUNDS - 180 Compounds are specifically claimed as (I), e.g. Inp-D-1Nal-D-Trp-3Pal-Lys-NH2 (Ia).

ADMINISTRATION - Dosage of (I) is 0.01-1000 mg/day and is administered orally, nasally, transdermally, transmucosally, intravenously, intraperitoneally, subcutaneously, topically or intramuscularly.

EXAMPLE - 3-Benzothienylalanine(D-Bal)-D-Trp-Phe-amino piperidinylcarboxylic acid(Apc)-Rink amide resin was treated with Fmoc-isonipecotic acid(Inp) (0.27 mmol) using N,N-diisopropylcarbodiimide (0.27 mmol) in 1-hydroxy-benzotriazole (HOBT) and dimethylformamide (DMF). The resulting product was treated with piperidine (20 %) in DMF. The peptide was treated with triisopropylsilane (8 %) in trifluoroacetic acid (1.5 ml) at room temperature for 2 hours to yield Inp-D-Bal-D-Trp-Phe-Apc-NH2.

DEFINITIONS - Preferred Definitions:
A1 = Apc or Inp;

A2 = S5 or D-Bip;

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S5 = D-Bal, D-1Nal or D-2Nal;
     A3 = S5 or D-Trp; and
     A4 = 3Pal, 4Pal, Pff, Phe, Pim, Taz, 2Thi or Thr(Bzl).
L116 ANSWER 3 OF 7 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
     2004-035048 [03]
                        WPIX
CR
     2004-071004 [07]
     C2004-011624
DNC
     Treating a hyperproliferative disease (e.g. cancer, psoriasis, adenoma or
     atherosclerosis) in a subject comprises administering a composition of a
     human lactoferrin alone or in combination with standard
     anti-cancer therapies.
DC
     BARSKY, R; PERICLE, F; PETRAK, K; VARADHACHARY, A; WANG,
TN
     Y; O'MALLEY, B
PΑ
     (BARS-I) BARSKY R; (PERI-I) PERICLE F; (PETR-I) PETRAK K; (VARA-I)
     VARADHACHARY A; (WANG-I) WANG Y; (OMAL-I) O'MALLEY B; (AGEN-N)
     AGENNIX INC
CYC
     104
                     A1 20031204 (200403) * EN
PΙ
     WO 2003099323
                                                51
                                                      A61K038-40
        RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR HU IE IT KE LS
            LU MC MW MZ NL OA PT RO SD SE SI SK SL SZ TR TZ UG ZM ZW
         W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK
            DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR
            KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NI NO NZ OM PH PL
            PT RO RU SC SD SE SG SK SL TJ TM TN TR TT TZ UA UG US UZ VC VN YU
            ZA ZM ZW
     US 2004009895
                     A1 20040115 (200406)
                                                      A61K038-40
                                                                      <--
     US 2004082504
                     A1 20040429 (200429)
                                                      A61K038-40
                                                                      <--
     AU 2003273182
                                                      A61K038-40
                     A1 20031212 (200443)
                                                                      <--
     EP 1507554
                     A1 20050223 (200515)
                                                                      <--
                                          FN
                                                      A61K038-40
         R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IT LI LT LU LV
            MC MK NL PT RO SE SI SK TR
ADT
    WO 2003099323 A1 WO 2003-US14789 20030509; US 2004009895 A1
     Provisional US 2002-379441P 20020510, Provisional US
     2002-379442P 20020510, Provisional US 2002-379474P 20020510
     . US 2003-434769 20030509; US 2004082504 A1 Provisional US
     2002-379441P 20020510, Provisional US 2002-379442P 20020510
     , Provisional US 2002-379474P 20020510, US 2003-435319 20030509;
     AU 2003273182 A1 AU 2003-273182 20030509; EP 1507554 A1 EP 2003-755357
     20030509, WO 2003-US14789 20030509
FDT
    AU 2003273182 Al Based on WO 2003099323; EP 1507554 Al Based on WO
     2003099323
                          20020510; US 2002-379441P
PRAI US 2002-379474P
     20020510; US 2002-379442P
                                    20020510; US
                       20030509; US 2003-435319
                                                      20030509
     2003-434769
IC
     ICM A61K038-40
     ICS A23L001-305
AB
     WO2003099323 A UPAB: 20050303
     NOVELTY - Treating a hyperproliferative disease comprises administering
     orally, intravenously or topically to a subject a human
     lactoferrin composition in an amount sufficient to provide an
     improvement in the hyperproliferative disease in the subject.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the
     following:
          (1) a method of enhancing a mucosal immune response in the
     gastrointestinal tract in a subject, comprising administering orally to
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(2) a method of reducing growth of a neoplasm in a subject, comprising administering orally to the subject a human **lactoferrin** composition in an amount to reduce the growth of the neoplasm in the

subject;

the subject a human lactoferrin;

(3) methods of enhancing a systemic or local immune response

following the step of administering intravenously or topically to the subject a lactoferrin composition; and

(4) methods of stimulating interleukin-18 or GFM-CSF in a subject, comprising administering to the subject the lactoferrin composition.

ACTIVITY - Cytostatic; Antirheumatic; Antiarthritic; Antiinflammatory; Osteopathic; Vasotropic; Antiarteriosclerotic; Antipsoriatic. No biological data given.

MECHANISM OF ACTION - Gene therapy.

USE - The methods are useful in treating malignant neoplasms (e.g. melanoma or leukemia) and other hyperproliferative diseases such as rheumatoid arthritis, inflammatory bowel disease, osteoarthritis, leiomyomas, adenomas, lipomas, hemangiomas, fibromas, vascular occlusion, restenosis, atherosclerosis, pre-neoplastic lesions, carcinoma in situ, oral hairy leukoplakia or psoriasis.

Dwq.0/5

FS CPI

FΑ AB: DCN

CPI: B04-N02; B14-C09A; B14-C09B; B14-E10C; B14-F01G; MC

B14-F07; B14-H01; B14-H01A; B14-H01B; B14-N17C

TECH

UPTX: 20040112 TECHNOLOGY FOCUS - BIOTECHNOLOGY - Preferred Method: In treating a hyperproliferative disease, the human lactoferrin composition is dispersed in a pharmaceutical carrier. The human lactoferrin is a recombinant human lactoferrin. The method further comprises administering an antacid in conjunction with the human lactoferrin composition. The hyperproliferative disease is cancer, which comprises a neoplasm: The neoplasm is selected from melanoma, non-small cell lung, small-cell lung, lung hepatocarcinoma, retinoblastoma, astrocytoma, glioblastoma, leukemia, neuroblastoma, squamous cell, head, neck, gum, tongue, breast, pancreatic, prostate, renal, bone, testicular, ovarian, mesothelioma, sarcoma, cervical, gastrointestinal, lymphoma, brain, colon, and bladder neoplasm. The neoplasm is a hematopoietic neoplasm selected from acute myelogenous leukemia, acute lymphoblastic leukemia, myelodysplastic syndrome, chronic myelomonocytic leukemia, juvenile myelomonocytic leukemia, multiple myeloma and chronic lymphocytic leukemia. The hyperproliferative disease is selected from rheumatoid arthritis, inflammatory bowel disease, osteoarthritis, leiomyomas, adenomas, lipomas, hemangiomas, fibromas, vascular occlusion, restenosis, atherosclerosis, pre-neoplastic lesions, carcinoma in situ, oral hairy leukoplakia, and psoriasis. Alternatively, the method comprises supplementing the mucosal, local or systemic immune system in a subject by increasing the amount of human lactoferrin in the gastrointestinal tract, in the vicinity of a tumor, or in the systemic circulation, respectively. The human lactoferrin stimulates the production of interleukin-18 or GM-CSF. Treating a hyperproliferative disease alternatively comprises administering to the subject the human lactoferrin composition in combination with chemotherapy, biotherapy, immunotherapy, surgery or radiotherapy. The lactoferrin may also be a bovine lactoferrin. The composition is a topical gel, a solution, capsule or a tablet having a lactoferrin concentration of about 0.01-20 (preferably 1.0-8.5)%. In enhancing a mucosal immune response in the gastrointestinal tract in a subject, the lactoferrin stimulates interleukin-18 or GM-CSF in the gastrointestinal tract. The interleukin-18 stimulates the production, maturation or activity of immune cells, such as T-lymphocytes or natural killer cells. The T-lymphocytes are selected from CD4+, CD8+ and CD3+ cells. The GM-CSF stimulates the production, maturation or activity of immune cells such as dendritic or other antigen presenting cells. The subject suffers from a hyperproliferative disease. The method further comprises additionally administering radiotherapy. Enhancing the mucosal or systemic immune response in the subject further comprises administering chemotherapy, immunotherapy, surgery, biotherapy, radiotherapy or their

combinations. The chemotherapy is a platinum-based agent (i.e. cisplatin) or a taxane-based agent (i.e. docetaxel). Reducing growth of a neoplasm in a subject further comprises administering the above-mentioned chemotherapy, immunotherapy, surgery, biotherapy, radiotherapy or their combinations. It further comprises administering radiotherapy.

UPTX: 20040112

ABEX

ADMINISTRATION - The amount of the composition that is administered orally is about 1 mg to about 100 g per day, preferably about 20 mg to about 10 g per day (claimed). When administered intravenously or topically, the amount of the composition is about 0.1 microg to about 10 g per day, preferably about 1 mug to 1 g per day (claimed). The composition may also be administered by intratumoral, nasal, buccal, rectal, vaginal, intramuscular, intraperitoneal, intraarterial or dermal means.

L116 ANSWER 4 OF 7 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN AN 2003-712670 [67] WPIX

DNC C2003-196034

Use of human apo-lactoferrin and peptides derivable from human lactoferrin for the production of composition useful for e.g. treating and preventing vascular disease.

DC B04

IN NORRBY, K

PA (NORR-I) NORRBY K

CYC 29

PI WO 2003072129 A1 20030904 (200367) * EN 14 A61K038-40 <-RW: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IT LU MC NL PT SE
SI SK TR

W: AU JP US

AU 2003210086 A1 20030909 (200428) A61K038-40 <--EP 1478387 A1 20041124 (200477) EN A61K038-40 <--

R: AT BE BG CH CY CZ DE DK EE ES FI FR GB GR HU IE IT LI LU MC NL PT SE SI SK TR

ADT WO 2003072129 A1 WO 2003-SE329 20030227; AU 2003210086 A1 AU 2003-210086 20030227; EP 1478387 A1 EP 2003-743090 20030227, WO 2003-SE329 20030227

FDT AU 2003210086 A1 Based on WO 2003072129; EP 1478387 A1 Based on WO 2003072129

PRAI SE 2002-598 20020227

IC ICM A61K038-40 ICS A61P009-00

AB WO2003072129 A UPAB: 20031017

NOVELTY - In the production of a composition, a substance containing human apo-lactoferrin and/or peptides derivable from human lactoferrin and/or its natural metabolites or equivalent analogs is used.

ACTIVITY - Antianginal; Cerebroprotective; Cardiant; Antiulcer; Antialopecia.

MECHANISM OF ACTION - VEGF165 induced angiogenesis inhibitor.

Lactoferrin, dissolved in saline, was given by tube feeding twice daily from Sunday afternoon (Day-1) to Friday afternoon (Day 4). Vehicle controls received saline by tube feeding. The angiogenesis treatment with VEGF was given intraperitoneally on Days 0 - 4 (twice daily). The results for test/control groups were vascularized area = 12.09 plus or minus 1.49/1.18 plus or minus 0.5, microvascular length = 1.465 plus or minus 0.077/0.28 plus or minus 0.04, and total microvascular length = 17.72 plus or minus 2.19/0.33 plus or minus 0.14 respectively. The results demonstrated that oral administration of apo-hLE significantly enhanced the VEGF mediated angiogenic response.

USE - For treating and/or preventing vascular disease and/or states of tissue hypoperfusion (including impending or manifested stroke, ischemic heart disease e.g. angina pectoris or impending or manifested myocardial infarction), or peripheral artery occlusive disease with or without impending gangrene and/or state of depressed VEGF induced angiogenesis associated with peptic ulcer, leg ulcer or local or

generalized hair loss) with hypoxia and/or ischemic consequences (claimed).

ADVANTAGE - The method is used in as an alternative to bypass surgery or any therapeutic angiogenesis options. Dwg.0/0

FS CPI

FA AB; DCN

MC CPI: B04-C01; B04-N02; B04-N02B; B14-E08; B14-F01;

B14-F01D; B14-F02; B14-F02D;

B14-F02D1; **B14-F02F**; B14-K01; B14-N16; B14-R02 UPTX: 20031017

TECH

TECHNOLOGY FOCUS - BIOLOGY - Preferred Substance: The substance is human apolactoferrin, human lactoferricin, a peptide constituted of 12 -40 amino acids of human lactoferrin counted from the N-terminal end or its modified version, a peptide formed of a sequences constituted of 16 - 40 amino acids and 18 - 40 amino acids from the N-terminal and of human lactoferrin or its modified version, a peptide corresponding to 18 - 31 residues of human lactoferrin (where C-20 is replaced by A, Q-22 is replaced by K and N-26 is replaced by D), a peptide formed of the amino acids in positions 12 - 31, counted from the N-terminal end, in the sequence constituting human lactoferrin, or its modification or its fragment consisting of at least 7 amino acids, a peptide consisting of 11 - 17 amino acids corresponding to the sequences that begin with one of the amino acids in positions 15 - 21 and end with the amino acid in position 31, counted from the N-terminal and, in the sequence constituting human lactoferrin or its modification, or a peptide consisting of 12 amino acids based on the sequence consisting of the amino acids in positions 20 - 31 in human lactoferrin, counted from the N-terminal end.

ABEX UPTX: 20031017

ADMINISTRATION - The route of administration is oral, parenteral, local or by inhalation. No dosage given. EXAMPLE - No relevant example given.

L116 ANSWER 5 OF 7 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

AN 2003-598327 [56] WPIX

DNC C2003-162395

TI Composition for e.g. improving lipid metabolism useful in treating lifestyle-related diseases like hypercholesterolemia and (severe) obesity comprises lactoferrin proteins and their enzymatic digestion products,.

DC B04

IN ANDO, K; HARADA, E; SHIMIZU, H; TAKEUCHI, T

PA (NUCL-N) NUCLEAR RECEPTOR LIGAND CO LTD; (NRLP-N) NRL PHARMA INC; (ANDO-I)
ANDO K; (HARA-I) HARADA E; (SHIM-I) SHIMIZU H; (TAKE-I) TAKEUCHI T
CYC 103

PI WO 2003057245 A1 20030717 (200356)* JA 51 A61K038-40 <-RW: AT BE BG CH CY CZ DE DK EA EE ES FI FR GB GH GM GR IE IT KE LS LU
MC MW MZ NL OA PT SD SE SI SK SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ OM PH PL PT RO RU SC SD SE SG SK SL TJ TM TN TR TT TZ UA UG US UZ VC VN YU ZA ZM ZW

AU 2002359949 A1 20030724 (200421) A61K038-40 <-EP 1466621 A1 20041013 (200467) EN A61K038-40 <--

R: AL AT BE BG CH CY CZ DE DK EE ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT RO SE SI SK TR

US 2005020484 A1 20050127 (200509) A61K038-40

ADT WO 2003057245 A1 WO 2002-JP13858 20021227; AU 2002359949 A1 AU 2002-359949 20021227; EP 1466621 A1 EP 2002-793463 20021227, WO 2002-JP13858 20021227; US 2005020484 A1 WO 2002-JP13858 20021227, US 2004-500245 20040625

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FDT AU 2002359949 A1 Based on WO 2003057245; EP 1466621 A1 Based on WO
     2003057245
PRAI JP 2001-400641
                          20011228
     ICM A61K038-40
     ICS A61K009-14; A61K009-144; A61K009-16; A61K009-166; A61K009-20;
          A61K009-200; A61K009-48; A61K009-488; A61K038-16; A61K038-166;
          A61P001-16; A61P001-166; A61P003-04; A61P003-044; A61P003-06
          ; A61P003-066; A61P003-10; A61P003-100; A61P009-12;
          A61P009-122
AB
     WO2003057245 A UPAB: 20030903
     NOVELTY - Composition comprises one or more of lactoferrin
     proteins (I) including lactoferrin and conalbumin, and their
     enzymatic digestion products such as lactoferrin and peptides of
     conalbumin corresponding to lactoferrin.
          DETAILED DESCRIPTION - An INDEPENDENT CLAIM is also included for the
     preparation of the composition comprising mixing (I) with
     pharmaceutically-acceptable additives in dry state for pressing into
     grains for filling capsules, or for producing powders, granules or
     tablets;
          ACTIVITY - Antilipemic; Anorectic; Antidiabetic; Hepatotropic.
          Test details are described, but no results are given.
          MECHANISM OF ACTION - None given in source material.
          USE - The compositions are used for improving lipid metabolism and
     elevating basal metabolic rate, which is useful in treating
     lifestyle-related diseases such as hypercholesterolemia,
     hypertriglycedemia, low-density lipoprotein hypercholesterolemia,
     high-density lipoprotein hypocholesterolemia, obesity, fatty liver,
     cholesterol cholelithiasis (all claimed), hyperlipemia and type II
     diabetes.
     Dwq.0/14
FS
     CPI
FA
     AB; DCN
     CPI: B04-N06; B14-D02A2; B14-E12; B14-F06; B14-N12;
MC
          B14-S04
ABEX
                    UPTX: 20030903
     ADMINISTRATION - Administration is oral, e.g. in the form of powders,
     granules, tablets, capsules, enteric preparations particularly with
     coating resistant to gastric juice but soluble in the small intestine or
     as drinks. The dosage is 0.1-50000 (preferably 10-2000) mg daily.
     EXAMPLE - A composition was formulated from lactoferrin (1 part)
     and potato starch (1 part) with water to produce grains, which were then
     filled into capsules at 150 mg each.
L116 ANSWER 6 OF 7 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN
     1994-307580 [38]
                        WPIX
                        DNC C1994-140034
DNN N1994-242039
     Denatured low density lipoprotein-combining agent - comprises
     lactoferrin and/or hydrolysate shows antimicrobial activity and is
     used in treatment of progressive arteriosclerosis.
DC
     B04 P34
     (MORG) MORINAGA MILK IND CO LTD
PΑ
CYC
PΤ
     JP 06234655
                     A 19940823 (199438)*
                                                      A61K037-14
                                                                      <--
     JP 3497195
                     B2 20040216 (200413)
                                                 8
ADT
     JP 06234655 A JP 1993-23055 19930210; JP 3497195 B2 JP
     1993-23055 19930210
FDT
     JP 3497195 B2 Previous Publ. JP 06234655
PRAI JP 1993-23055
                          19930210
     ICM A61K037-14; A61K038-16
     ICS A61K037-18; A61K038-00; A61M001-36; A61P009-10
     JP 06234655 A UPAB: 19941115
AB
```

Denatured low density lipoprotein-combining agent comprises (a)

lactoferrin and/or (b) hydrolysate.

USE/ADVANTAGE - Lactoferrin is iron-combining protein derived from milk, tear, saliva, blood etc. and is known to show antimicrobial activity and to involve in inflammation. It was found to specifically combine with denatured LDL e.g. LDL oxidised with free radical, selectively remove the LDL, and inhibit incorporation of the LDL into macrophage so that foaming of macrophage is inhibited. This agent is therefore useful for prevention and treatment of progressive arteriosclerosis.

Dwg.0/6

FS CPI GMPI

FA AB; GI

MC CPI: B04-N02; B14-F07

L116 ANSWER 7 OF 7 WPIX COPYRIGHT 2005 THE THOMSON CORP on STN

AN 1994-238662 [29] WPIX

DNC C1994-108974

TI Brain protectant for preventing ischaemic diseases without side effects - comprising 31 specified peptide(s), prepared by **lactoferrin** hydrolysis.

DC B04

PA (MORG) MORINAGA MILK IND CO LTD

CYC 1

PI JP 06172200 A 19940621 (199429)* 11 A61K037-02 <--

ADT JP 06172200 A JP 1992-327738 19921208

PRAI JP 1992-327738 19921208

IC ICM A61K037-02

ICA A61K037-14; A61K037-18; C07K005-08; C07K005-10; C07K007-06; C07K007-08; C07K007-10

ICI C07K099:

AB JP 06172200 A UPAB: 19940907

Compsn. comprises 31 specified peptides, their derivs. or salts, or their mixts..

Lactoferrin is pref. chemically or enzymically hydrolysed to give 31 peptides having 3-47 amino acid sequences with brain protecting property. The peptides also have antimicrobial activity and their pharmaceutical prepns. may be prepared without addition of antiseptics. The peptides are administered at least at doses of 10 mg for parenteral and 100mg for oral admin.

USE/ADVANTAGE - Stable, heat resistant, water soluble and antimicrobial brain protectant is used for the prevention of ischaemic diseases without side effects.

In an example, a peptide having 25 amino acids, Phe-Lys-Cys-Arg-Arg-Trp-Gln-Trp-Arg-Met -Lys-Lys-Leu-Gly-Ala-Pro-Ser -Ile-Thr-Cys-Val-Arg-Arg-Ala-Phe, exhibited brain protecting effect in decapitated male ddy mice at doses of at 10 mg/kg by intraperitoneal, and 50 mg/kg by subcutaneous admin. respectively.

Dwg.0/0

FS CPI

FA AB; GI; DCN

MC CPI: B04-C01; B04-N02A; B14-A01; B14-F02D1

=> d his

(FILE 'HOME' ENTERED AT 12:52:20 ON 09 MAR 2005) SET COST OFF

FILE 'REGISTRY' ENTERED AT 12:52:25 ON 09 MAR 2005 E LACTOFERRIN

L1 224 S E3, E4, E8

L2 23 S E1, E2, E5-E7 NOT L1

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FILE 'HCAPLUS' ENTERED AT 12:53:57 ON 09 MAR 2005
                E LACTOFERRIN/CT
L3
           3804 S E6-E10
                E E6+ALL
L4
           3828 S E4,E3
           4954 S LACTOFERRIN OR LACTOTRANSFERRIN
L5
L6
            349 S L1 OR L2
L7
           5056 S L3-L6
              1 S LACTO() (FERRIN OR TRANSFERRIN OR TRANS FERRIN)
L8
L9
           5056 S L7,L8
                E ATHEROSCLEROSIS/CT
L10
          26376 S E3, E4
                E E3+ALL
L11
           5033 S E10-E13
          45035 S E9,E11,E12,E13/BI
L12
                E E8+ALL
           8095 S E8
L13
L14
          10796 S E8/BI
                E E15+ALL
           8237 S E4
L15
             32 S L9 AND L10-L15
L16
                E CARDIOVASCULAR/CT
                E E5+ALL
         63843 S E3+NT
L17
                E E19+ALL
L18
         245711 S E4, E3+NT
                E E250+ALL
L19
         375833 S E3+NT
                E HEART DISEASE/CT
                E E4+ALL
                E E2+ALL
L20
         82991 S E8, E9, E7+NT
                E E92+ALL
L21
         216429 S E5, E4+NT
L22
          6523 S E10+OLD,NT
            189 S L9 AND L17-L22
L23
             12 S L9 AND CARDIOVASCULAR(L) (DISEASE OR DISORDER OR DYSFUNCTION?)
L24
L25
            194 S L16, L23, L24
     FILE 'REGISTRY' ENTERED AT 12:59:21 ON 09 MAR 2005
              1 S CHOLESTEROL/CN
L26
                E C-REACTIVE PROTEIN/CN
L27
              1 S E3
            106 S C REACTIVE PROTEIN
L28
     FILE 'HCAPLUS' ENTERED AT 13:00:01 ON 09 MAR 2005
L29
         107734 S L26 OR L27 OR L28
L30
             60 S L29 AND L9
            137 S (?CHOLESTER? OR CRP OR C REACTIVE(L) PROTEIN) AND L9
L31
L32
             30 S TRIGLYCER? AND L9
L33
             33 S ?VASCUL?(L)?INFLAM? AND L9
L34
              1 S ?VASCUL? (L) ?SPASM? AND L9
              1 S BLOOD VESSEL+OLD, NT/CT (L) SPASM? AND L9
L35
             1 S BLOOD VESSEL, DISEASE+OLD, NT/CT (L) SPASM? AND L9
L36
             34 S PROTEIN?/CW,CT (L) C REACTIVE AND L9
L37
L38
              1 S BLOOD VESSEL, DISEASE+OLD, NT/CT (L) HYPERREACT? AND L9
              1 S BLOOD VESSEL+OLD, NT/CT (L) HYPERREACT? AND L9
L39
             6 S BLOOD VESSEL+OLD, NT/CT (L) SMOOTH MUSCL? AND L9
L40
             4 S BLOOD VESSEL, DISEASE+OLD, NT/CT (L) SMOOTH MUSCL? AND L9
L41
             6 S INFLAMM?/CW,CT (L) VASCUL? AND L9
L42
L43
             0 S INFLAMM?/CW,CT (L) PRO(L)CYTOKIN? AND L9
            10 S INFLAMM?/CW,CT (L) CYTOKIN? AND L9
L44
            37 S CYTOKINE?/CW,CT (L) ?INFLAM? AND L9
L45
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E CYTOKINE/CT
L46
             72 S E77+OLD, NT (L) ?INFLAM? AND L9
             14 S BLOOD VESSEL, DISEASE+OLD, NT/CT (L) ENDOTHEL? AND L9
L47
L48
             39 S BLOOD VESSEL+OLD, NT/CT (L) ENDOTHEL? AND L9
             12 S ENDOTHELIUM+OLD, NT/CT (L) VASCUL? AND L9
L49
                E HYPERCHOLESTEROL/CT
              7 S E5, E6 AND L9
L50
                E E5+ALL
L51
              0 S E5 AND L9
                E HYPERTRIGLYCER/CT
                E E4+ALL
              4 S E4, E5 AND L9
L52
                E LOW DENSITY LIPOPROTEIN/CT
                 E L DENSITY LIPOPROTEIN/CT
                E LIPOPROTEIN/CT
L53
             12 S E100-E109, E113, E114 AND L9
L54
             54 S E135-E146 AND L9
                E E51+ALL
L55
             56 S E2+NT (L) (LOW OR VERY LOW) () (DENSITY OR D OR DEN) AND L9
L56
             12 S E2+NT (L) HIGH() (DENSITY OR D OR DEN) AND L9
L57
             19 S E2+NT (L) (LDL OR VLDL OR HDL OR VHDL) AND L9
L58
            401 S L30-L57, L25
L59
              1 S US20040152623/PN OR WO2003-US38540/AP, PRN
                E VARADHACHARY A/AU
L60
             19 S E3, E7
                E GLYNN P/AU
L61
             53 S E3-E9, E17-E19
                E WANG Y/AU
L62
           2479 S E3, E40-E43
                E WANG YEN/AU
L63
             11 S E3, E34
             13 S E50
L64
                E ENGELMAYER J/AU
L65
              9 S E4
                E AGENNIX/AP,CS
                E AGENNIX/PA,CS
                E AGENIX/PA,CS
L66
             17 S E3-E21
L67
             10 S L59-L66 AND L58
             10 S L67 AND L3-L25, L29-L67
L68
L69
            341 S L58 AND (PD<=20021204 OR PRD<=20021204 OR AD<=20021204)
L70
            106 S L69 AND (PHARMACEUT? OR PHARMACOL?)/SC,SX
                E DRUG DELIVERY/CT
L71
              2 S E27-E31,E39 AND L70
L72
              0 S E53, E55, E58, E64, E70, E71 AND L70
L73
              0 S E89,E107 AND L70
L74
             50 S E6-E217 AND L70
                E E6+ALL
L75
              6 S E3-E5 AND L70
L76
             53 S E2+NT AND L70
L77
             53 S L71, L74-L76
L78
            106 S L70, L77
     FILE 'REGISTRY' ENTERED AT 13:36:48 ON 09 MAR 2005
     FILE 'REGISTRY' ENTERED AT 13:37:26 ON 09 MAR 2005
L79
             14 S 59-67-6 OR 943-45-3 OR 11041-12-6 OR 50925-79-6 OR 182815-43-
L80
            717 S (59-67-6 OR 943-45-3 OR 11041-12-6 OR 50925-79-6 OR 182815-43
L81
              1 S 9028-35-7
     FILE 'HCAPLUS' ENTERED AT 13:38:11 ON 09 MAR 2005
L82
              6 S L79, L80, L81 AND L78
L83
              8 S L79, L80, L81 AND L69
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L84
             1 S BILE ACID (L) SEQUESTR? AND L69, L78
             7 S L82-L84, L78 AND ?ATHEROSCLERO?
L85
L86
             23 S L69 AND ?ATHEROSCLERO?
            23 S L85, L86
L87
               SEL DN AN 1 7 12
L88
             3 S L87 AND E1-E7
L89
             99 S L78 NOT L87
               SEL DN AN 8 19 21 68 89
               DEL SEL
               SEL DN AN 8 19 21 68 96
L90
             5 S L89 AND E1-E15
L91
             16 S L88, L90, L68
L92
             16 S L91 AND L3-L25, L29-L78, L82-L91
    FILE 'HCAPLUS' ENTERED AT 13:53:44 ON 09 MAR 2005
    FILE 'WPIX' ENTERED AT 13:55:12 ON 09 MAR 2005
L93
           1 S L59
L94
           161 S A61K038-40/IPC
L95
           693 S L5/BIX OR L8/BIX
           683 S ?LACTOFERRIN?/BIX
L96
L97
           752 S L94-L96
             4 S L97 AND (B14-D02A2 OR C14-D02A2 OR B12-H03 OR C12-H03)/MC
L98
L99
            80 S L97 AND (B14-F? OR C14-F? OR B12-F? OR C12-F?)/MC
            9 S L97 AND A61P009/IPC
L100
             2 S L97 AND A61P003-06/IPC
L101
            55 S L97 AND (P5? OR P814)/M0,M1,M2,M3,M4,M5,M6
L102
            97 S L98-L102
L103
            90 S L103 AND (PY<=2002 OR PRY<=2002 OR AY<=2002)
L104
            20 S L104 AND N135/M0, M1, M2, M3, M4, M5, M6
L105
            33 S L104 AND D05-H?/MC
L106
L107
            35 S L105, L106
               SEL DN AN 2
             1 S L107 AND E16-E17
L108
L109
             55 S L104 NOT L107
               SEL DN AN 10 15 21 23 52 54
             6 S L109 AND E18-E30
L110
             7 S L93,L108,L110
L111
            13 S L97 AND (ENGELMAYER ? OR GLYNN ? OR VARADHACHARY ? OR WANG ?)
L112
            16 S L97 AND AGEN?/PA
L113
            19 S L112,L113
L114
             2 S L111 AND L114
L115
             7 S L111,L115
L116
             17 S L114 NOT L116
L117
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FILE 'WPIX' ENTERED AT 14:30:50 ON 09 MAR 2005

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